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(54) Title: GENETIC ANALYSIS OF PEYER'S PATCHES AND M CELLS AND METHODS AND COMPOSITIONS TAR-
GETING PEYER'S PATCHES AND M CELL RECEPTORS

(57) Abstract: Methods of increasing of or decreasing the levels of a protein in a PP cell; methods of increasing antigen, vaccine,
DNA vaccine delivery to M cells, use of human serum albumin and other transport enhancing proteins to enhance oral drug delivery;
use of calreticulin to enhance oral antigen delivery, use of other cell surface proteins, receptors, and transporters to enhance delivery
to M cells of antigens or vaccine delivery vehicles, use of other cytoplasmic proteins to regulate intracellular trafficking and delivery
to mucosal immune sampling and processing systems.

TITLE OF THE INVENTION

5 GENETIC ANALYSIS OF PEYER'S PATCHES AND M CELLS AND METHODS AND
COMPOSITIONS TARGETING PEYER'S PATCHES AND M CELL RECEPTORS

CROSS REFERENCE TO RELATED APPLICATIONS

10 This application claims the benefit of U.S. provisional application 60/281,387 filed April 4,
2001, and U.S. provisional application 60/302,591 filed July 2, 2001.

FIELD OF THE INVENTION

15 This invention relates to the genetic analysis of M cells and methods and
compositions targeting M cell receptors.

BACKGROUND OF THE INVENTION

20 The Peyer's patch of the intestinal lining is a specialized tissue that allows the
immune system to identify foreign antigens that require an immune response. It is also a
potential pathway for orally delivered drugs to cross the intestinal barrier into the
bloodstream. Central to these properties are M cells, which populate the patch's epithelial
sheet. In view of the importance of the Peyer's patch and its M cells for the immune
response and drug delivery, it is desirable to identify the cell proteins important for these
phenomena. It is also desirable to increase the amounts of such important proteins in order
25 to either facilitate the immune response and drug delivery or promote the conversion of non-
M cells to M cells.

30 Similarly, it is important to identify and further decrease the levels of proteins whose
absence or down-regulation in expression facilitates the immune response and drug delivery,
or promotes the conversion of non-M cells to M cells.

BRIEF SUMMARY OF THE INVENTION

Increasing the levels of a protein or antigen-protein combination

In a first general aspect, the invention is a method of increasing the levels of a protein

in a Peyer's patch cell, said method comprising delivering to said cell a nucleic acid coding for a protein, wherein absent said increase, the levels of said protein or its mRNA is greater than in a non-Peyer's patch cell.

5 Peyer's patch cells of particular interest are M cells. The levels of a protein or its mRNA in Caco-2 cells co-cultured with Raji B cells are considered herein to be representative of such levels in a human Peyer's patch M cell. Monoculture Caco-2 cells are considered herein to be an appropriate non-Peyer's patch cell for purposes of comparison of such protein or mRNA levels.

10 The levels of a protein or its mRNA in rat Peyer's patch epithelial cells can be compared to their respective levels in a culture of rat normal gut epithelial cells. Absent evidence to the contrary, results of rat cells are assumed to be predictive of the results in human cells.

15 The presence of Increased levels of an mRNA, and therefore presumptively its protein, are indicated in the Table 2 and 3 by a **, a *, or an expression Fold Change greater than 1.00. Preferred are those indicated by a ** or an expression Fold Change greater than 2.00. Most highly preferred are those indicated by a **. The presence of decreased levels of an mRNA, and presumptively its protein, are indicated by a minus sign (-) or an expression Fold Change less than 1.00. Preferred targets are those indicated by a minus sign or an expression Fold Change less than 0.50.

20 In embodiments of particular interest, the protein is a receptor, a transporter, cell surface antigen, or cell adhesion molecule, especially a receptor. In other embodiments of particular interest, the protein is selected from the group consisting of nucleoside diphosphate kinases and member of the 14-3-3 family.

25 In the methods of greatest interest, the nucleic acid is delivered to a human cell. There are many delivery options, one of which is to deliver it by the oral route with the cell in a human, another to deliver it to a cell outside a human.

30 In an important variation of the method, a nucleic acid coding for a tumor antigen or foreign peptide is also delivered to the Peyer's patch cell. The purpose of this aspect of the invention is to improve the immune response to a tumor antigen or the foreign peptide. Normally, therefore, the foreign peptide will be that of a virus or infectious microorganism. A tumor antigen is one that is more abundant in a tumor cell than its normal counterpart.

Decreasing the levels of a protein

Another general aspect of the invention is a method of decreasing the levels of a

protein in a Peyer's patch cell, said method comprising delivering to said cell an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference (RNAi) nucleic acid molecule, said anti-sense, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for said protein, wherein absent said anti-sense nucleic acid molecule, ribozyme or RNAi nucleic acid, the levels of said protein or its mRNA are less than in a non-Peyer's patch cell. More preferably the anti-sense nucleic acid is complementary to a sequence of at least 15 nucleotides of the mRNA of the protein, and most preferably to a sequence of at least 30 nucleotides of the mRNA of the protein. It is preferred that the protein is coded for by a gene with an expression Fold Change denoted by a minus sign (-) or an expression Fold Change less than 0.50.

In a particular embodiment, the latter method comprises delivering to said cell an anti-sense nucleic acid molecules, a ribozyme or RNAi nucleic acid molecules, said anti-sense, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for at least 5 different proteins, wherein absent said anti-sense, ribozyme or RNAi nucleic acid molecule, the levels of each of said proteins or its mRNA are less than in a non-Peyer's patch cell.

Alternatively described, the latter invention is a method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell an anti-sense nucleic acid molecule, ribozyme or RNAi nucleic acid molecules, said anti-sense, ribozyme or RNAi nucleic acid forming a double-stranded molecule with part or all of the mRNA for said protein, wherein absent said anti-sense, ribozyme or RNAi nucleic acid molecule, the levels of said protein or its mRNA are less than in a non-Peyer's patch cell.

Cells of the invention

A human or rat cell to which any of the above methods in this Brief Summary of the Invention section has been applied, or the progeny of said cell, is also an aspect of the present invention.

Delivery enhancement using a targeting ligand which targets a receptor, a transporter or a cell-surface molecule expressed on surface of M cells or Peyer's patch tissue cells

In another general aspect, the invention is a method of targeting an antigen or a drug delivery vehicle containing an antigen, or a drug delivery vehicle containing an antigen and adjuvant, or a drug delivery vehicle containing a drug, or a viral vector, or a bacterio-phage vector such as, but without limitation M13 or Fd, or a bacterial vector or a gene delivery

vector expressing an antigen of interest, or a viral vector, or a bacterio-phage vector such as, but without limitation M13 or Fd, or a bacterial vector or a gene delivery vector expressing a gene product(s) to M cells of Peyer's patch tissue, by targeted delivery to receptors, or to transporters or to other cell surface proteins which are found to be expressed on the cell surface of M cells or other cells found within Peyer's patch tissue, or which are found to be differentially expressed on these cells. Said gene product(s) coded by the viral vector, or a bacterio-phage vector such as, but without limitation M13 or Fd, or a bacterial vector or a gene delivery vector regulate the function of Peyer's patch cells to M cell phenotype or regulate M cell function to increase their immuno-surveillance or antigen presentation to the mucosal immune system.

In one embodiment, a phage display library such as M13 or Fd which express random peptide sequences on the surface of the phage, coded by example gene III or gene VII of M13 or Fd bacteriophage, can be screened by in vivo panning against example Peyer's patch tissue found in vivo in the GIT, in order to discover and identify phage or targeting ligands which specifically target M cells or Peyer's patch tissue in vivo in the GIT; such phage which target M cells and Peyer's patch tissue can subsequently be genetically engineered to encode a gene or genes of interest such as a DNA vaccine gene, a gene coding for an antigen of interest together with gene(s) which modify M cell function and which enhance the immuno-responsiveness of the M cells to the antigen or DNA vaccine product coded by the genetically engineered bacteriophage genome.

Delivery enhancement using transport enhancing proteins

Another invention disclosed herein is a method for enhancing transport of a drug through the gastrointestinal tract, said method comprising orally administering said drug in a composition that comprises a transport-enhancing protein, said transport-enhancing protein selected from the group consisting of human serum albumin (HSA), clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, and Ca^{2+} -dependant phospholipase A_2 (Ca^{2+} pl a_2), or a homolog that has at least 80% amino acid identity with said transport-enhancing protein over a length of said transport-enhancing protein identical to the homolog. In a preferred embodiment, the homolog has at least 90% amino acid identity with the transport-enhancing protein over a length of the transport-enhancing protein identical to the homolog. In a more preferred embodiment, the transport-enhancing protein is selected from the group consisting of human serum albumin (HSA), clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, and Ca^{2+} pl a_2 .

Method of delivering a vaccine to a target cell

Further invention disclosed herein is a method of delivering a vaccine to a target cell, said method comprising utilizing as the target cell a Peyer's patch cell in which a protein or mRNA is upregulated.

5

Method of decreasing the levels of a protein

Yet, another invention disclosed herein is a method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell a DNA molecule coding for an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule (RNAi), said anti-sense molecule, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for said protein, wherein absent said anti-sense molecule, ribozyme or RNAi nucleic acid, the levels of said protein or its mRNA is less than in a non-Peyer's patch cell.

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Method of increasing the extent to which the function of a protein is carried out

Another invention disclosed herein is a method of increasing the extent to which the function of a protein is carried out in a Peyer's patch cell, said method comprising delivering to said cell a nucleic acid coding for said protein, wherein absent said delivery, the level of said protein or its mRNA is greater in said cell than in a non-Peyer's patch cell.

20

Chimeric protein that comprises two or more segments, each of said segments enhancing a different step in the peptide transport process

Another invention disclosed herein is a chimeric protein that comprises two or more segments, each of said segments enhancing a different step in the peptide transport process, said steps selected from the group consisting of binding to a cell such as an M cell, transporting the peptide into the cell such as an M cell, presenting the chimeric protein to a protein processing pathway within a cell such as an M cell in order to maximise processing in a way to optimize presentation of the processed chimeric peptides to epitopes suitable for immune activation, transporting the peptide through the cell such as an M cell, and transporting the peptide out of the cell such as an M cell to an underlying immune cell such as a B-cell or T-cell.

25

30

Delivery enhancement using calreticulin and other proteins

Another method disclosed herein is a method to facilitate intracellular trafficking of an antigen that has been orally delivered by itself or as part of a composition or particle, said method comprising administering calreticulin.

5 Related to the latter invention is a chimeric protein comprising the amino acid sequences for (1) calreticulin, rab family proteins and and/or a ribosomal protein, and (2) a second polypeptide. Also related is a method of administering a polypeptide, where said polypeptide is part of the chimeric protein and wherein said chimeric protein is orally administered.

10

DETAILED DESCRIPTION OF THE INVENTION

The present invention and the related research were intended to improve targeted vaccine delivery and targeted gene delivery methods, especially as they relate to Peyer's patch cells. In significant part, this was achieved by identifying proteins whose up-regulation or down-regulation would indicate their possible or probable role in cellular functions important to vaccine and or drug delivery. In some cases, such as receptors, the proteins are important from the point of view of cell specificity during the delivery process. In many cases, the proteins have functions that are important after the vaccine or drug enter the cell.

20 Closely related to those inventions and research goals, was the concept that in M cells there would be proteins that, as compared to M cell precursors, were up-regulated or down-regulated. The identification of such proteins provides a strategy for altering M cell precursors so as to shift their phenotype toward that of M cells.

25 As indicated, one aim of the research related to the present invention was to determine if there were detectable differences in protein/gene expression between: (1) Peyer's patch (PP) and non-Peyer's patch (NPP) rat gastrointestinal tract (GIT) tissue and (2) M cell enriched follicle-associated epithelium of Peyer's patch (PP FAE) tissue. This was done with a view to finding novel or highly expressed ligand targeting sites on the Peyer's patch or M cells as well as other protein relevant to the delivery of drugs across the GIT.

30 This invention is based in part on the discovery of over-expression of a range of genes in Peyer's patch (PP) tissue from rat small intestine in comparison to normal non-Peyer's patch (NPP) small intestine tissue.

This invention is also based on the discovery of over-expressed genes in co-cultures of Caco-2 cells. The idea was to use genetic mapping of the M cell co-culture, e.g. Caco-2

cells co-cultured with Raji cells versus a monolayer of Caco-2 cells, to ascertain the differences in epithelial gene expression between M cells and enterocytes. It became immediately apparent that some of these gene products are going to be unique apical membrane proteins (e.g. receptors, transporters, adhesion proteins) in M cells. By
5 examining the differences between M cells and enterocytes in vitro and in vivo, one could identify key targets that can be used to generate M cell specific ligands. These ligands can then be used for targeting oral vaccines in particles.

The identification of over-expressed ribosomal proteins or homologues/related proteins thereof indicates a generally higher protein turnover or protein synthesis capacity
10 in PPs or a possible role for such ribosomal proteins (or homologues thereof) in other cellular functions such as protein chaperoning, endocytosis, trafficking of proteins/antigens/particulates/viruses uptaken from the lumen of the gastrointestinal tract (GIT) and/or from the M cells to underlying immune cells, antigen presenting cells, dendritic cells, B cells, other cell types.

The identification of a series of transcription factors (TFs) that are over-expressed in PP tissue versus the control enterocyte GIT tissue is considered herein to indicate a role for
15 such TFs in the development of M cell phenotype, in conferring M cell phenotype and/or in programming M cells to prime other downstream cellular events leading to a better or more efficacious immune outcome following antigen presentation. The co-delivery of genes
20 coding for such TFs with either antigens themselves and /or with gene(s) coding for antigen(s) of question to M cells and/or PP tissue following oral administration provides the basis for a more efficacious and pronounced immune outcome when the TF coding genes are key or vital for driving M cells / PP tissue to an effective immune outcome.

The general over-expression of a number of proteins species in PPs versus NPPs, both membrane and cytosolic-associated was also determined by a novel technique of
25 enrichment and M cell selection following enrichment of the follicle-associated epithelium (FAE) of Peyer's patch (PP FAE) by ethylene-diamine tetra-acetic acid (EDTA) extraction and recovery of M cells / PP FAE. Such novel or differentially expressed proteins have significant implications for the use of this protein expression information and methods of
30 selection / enrichment of M cells / PP tissue for the targeting of drug/vaccine uptake to Peyer's patches. Among proteins found to be over expressed in rat PP tissue following this enrichment technique was the human serum albumin homolog which is considered here to have implications for drug / cargo transport from the GIT either into or across intestinal tissue including PP tissue and systemic delivery of same to the blood.

Incorporation by reference

All references cited herein are incorporated herein by reference in their entireties.

5 All GenBank records specified by their accession numbers are incorporated herein by their entireties.

The GenBank amino acid sequences and nucleotide sequences specified by their GenBank ID number are incorporated by reference herein. All GenBank records corresponding to those ID numbers are incorporated herein in their entirety. Absent a date specifying the date of the record, the date of the record is the filing date of this application.

10 Many of GenBank sequences specified by their GenBank ID numbers are reproduced herein in the section "Amino acid sequences and nucleotide sequences corresponding to selected GenBank ID numbers." The CDS line refers to the exon(s).

15 Any GenBank ID numbers specified herein, absent a decimal point and an integer following that decimal point, is for GenBank version 1 of that sequence. Any GenBank ID number that has a decimal point and an integer following it is the GenBank version number.

20 The invention will be illustrated in more details with reference to the following Examples, but it should be understood that the present invention is not deemed to be limited thereto.

EXAMPLESExample 15 Preparation of cytosolic (S100) and membrane (P100) proteins from rat PP and NPP tissues

Protein samples were prepared from PP and NPP tissue extracted from freshly sacrificed rats. These protein samples underwent electrophoresis on denatured SDS-PAGE gels and were stained using two different standard proteins Commassie Blue stains. Subsequently, fresh PP and NPP tissue samples were fractionated into cytosolic (S100) and
10 membrane (P100) proteins and these samples were also electrophoresed on SDS-PAGE in order to compare S100 and P100 fractions in both PP and NPP tissues.

Example 215 Preparation of GIT tissue or co-culture cell membrane (P100) and cytosolic (S100) fractions

The fractions were prepared using the following procedure:

1. Scrape the co-culture cells into PBS and pool cells into a universal.
2. Centrifuge the cells for 5 minutes at 1,500 rpm.
3. Remove the supernatant.
- 20 4. Re-suspend the cell pellet in 3 volumes of ice-cold HED buffer, and allow it to swell for 5 minutes on ice.
5. Homogenize the cells for 30 seconds.
6. Centrifuge the homogenate in hard walled tubes at 40,000rpm for 45minutes at 4°C in a Beckmann Ultra Centrifuge (rotor Ti90).
- 25 7. Remove the supernatant (S100) and re-suspend the pellet (P100) in 3 volumes of HEDG buffer, before centrifugation again at 1000rpm for 2min. Remove the supernatant and store on ice. Repeat the procedure and add the second supernatant to the first.
8. Determine the protein concentration (using the Bio-Rad protein assay).
9. All fractions were stored at -80°C.

30 The following reagents were used in the above methods:

HED buffer (20mM HEPES pH 7.67), 1mM EGTA, 0.5mM dithiothreitol, 1mM phenylmethylsulphonyl fluoride (PMSF):

HEPES (pH to 7.67)	0.5206g
EGTA	38.04mg
35 Dithiothreitol	7.71 mg

Distilled water to 100ml

10µl PMSF stock solution was added to 1ml of buffer prior to use.

HEDG buffer (the same as HED buffer plus 100mM NaCl, 10% glycerol)

5	NaCl	0.584g
	Glycerol	11.4ml
	HEPES (pH to 7.67)	0.5206g
	EGTA	38.04mg
	Dithiothreitol	7.71 mg

10	Distilled water	to 100ml
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10µl PMSF stock solution was added to 1ml of buffer prior to use.

PMSF(100mM) stock solution

	PMSF	17.42mg
15	Isopropanol	

Example 3

Isolation of epithelial sheaths from rat Peyer's patch and non-Peyer's patch tissue

20 The M cell is a very elusive cell type, at least in terms of isolating a purified population. Previous attempts have found that when M cells are separated and purified and put into culture they very quickly lose their characteristic morphology and probably gene/protein expression profile. In many cases this is due to the length of time taken to isolate and purify the cells from the very homogenous mix of cells in a Peyer's patch. We

25 desired a quick and routine method to enrich for M cells in Peyer's patch samples. M cells are only contained in the epithelium of Peyer's patches, the so-called follicle associated epithelium (FAE), while underneath the epithelial layer lays all the B and T lymphocytes, dendritic cells etc. So by isolating the epithelium away from the rest of the Peyer's patch

30 dome, we are greatly enriching it for the M cell population. Previously, treatment of mouse intestinal tissue with EDTA was shown to cause separation of the epithelium as a sheet from the rest of the tissue, allowing for it's specific isolation (Bjerknes M and Cheng H (1981). Methods for the isolation of intact epithelium from the mouse intestine. *Anat. Rec.*, (199):565). This method was adapted for the isolation of FAE from rat Peyer's patch. Control epithelium from normal gut tissue (no Peyer's patches) was used as a control.

Epithelial sheaths were prepared using EDTA method comprising the following steps:

1. Sacrifice the rats (Wistar) by cervical dislocation.
2. Remove the entire length of the GIT tract from (but not including) the stomach to the caecum and place in a dish of PBS (at room temperature).
3. Excise the Peyer's patches, taking care to remove as much normal non-PP GI tissue as is visible. Rinse briefly in PBS.
4. Also take samples of normal non-Peyer's (NPP) tissue close to the patches, rinse in PBS and treat as for the PPs (steps 5-9, 11-12).
5. Pool the PP's from the entire GI section in Hank's Buffered Saline Solution (HBSS, Gibco Life Sciences) with 0.011M glucose and 25mM Hepes.
6. When pooling is complete, place PP sections into 15-20ml of HBSS (with 0.011M glucose and 25mM Hepes) along with 40mM of EDTA into a small conical flask.
7. Add a stirrer bar to the flask, place on a stirring plate and spin the PP's for 15 min at RT.
8. After 15 minutes pipette the PP solution vigorously with a wide-bore 3ml plastic pasteur pipette.
9. Strain the supernatant through a 100micron nylon cell strainer (from FALCON™, 352360).
10. Move the filter to another 50ml tube and wash out the residue material on the filter with HBSS (with 0.011M glucose and 25mM Hepes, no EDTA). This residue contains the majority of the PP dome epithelial sheaths.
11. Centrifuge the PP residue material at 3000 rpm for 5min. Also centrifuge the NPP tissue supernatant at 3000 rpm for 5min.
12. Snap freeze the cell pellets and store at -70°C.

Example 4

Identification of over-expressed proteins in enriched M cells / PP FAE cells

Epithelial cell layers of Wistar rat PP (representing enriched M cells / PP FAE cells) and normal villi were extracted using EDTA as described in Example 3 above. Epithelial layers from numerous Patches and rats were pooled and the protein isolated into either cytosolic and membrane fractions following centrifugal separation. 2D gel electrophoresis (between isoelectric points pH 3.5 to 10) was performed on 50 µg of each fraction, wherein the gels were silver stained. The gels were overlaid, and numerous differentially expressed

proteins between the membrane fractions of PP and normal villi epithelia were observed. Further, protein samples underwent a second 2D gel electrophoresis, this time the gel was stained with a "special" silver stain, that did not inhibit the mass spectrometry analysis of individual spots. The differentially expressed proteins were identified and highlighted by gel overlay.

Thirty-seven protein spots were identified that were increased in PP over villi epithelial membrane fractions. Of these, 16 spots (the most highly over-expressed) were chosen for mass spectrometry analysis. The spots were digested with endoproteinase Lys-C/trypsin (8:1 ratio) and analyzed on a MALDI-MS. The spots, however, gave very poor spectra and only 4 of the 16 were identifiable. These were:

serum albumin;
calreticulin;
14-3-3 zeta (tentative ID; mouse); and
nucleoside diphosphate kinase B.

One protein showed homology to human serum albumin (HSA). Work by A. Fasano at Maryland had suggested that Zonulin, the human homologue of ZOT, showed sequence homology to human serum albumin (85% homology across the limited sequence available from the Fasano's work). Given our finding that a protein differentially expressed in rat PP tissue shows homology to HSA, we propose that HSA (or a homologue or splice variant thereof) is involved in drug transport in the GIT, in particular Peyer's patch tissue of the GIT.

Calreticulin is a 46-kDa Ca (2+)-binding chaperone of the endoplasmic reticulum membranes. This protein binds Ca (2+) with high capacity, affects intracellular Ca (2+) homeostasis, and functions as a lectin-like chaperone. Given the over-abundance of expression of this protein in epithelial layers selected from PP tissue and the role of this protein as a lectin-like chaperone, we propose that this protein is a valuable protein target to aid or facilitate the intracellular trafficking of antigens or antigens in particles following targeted delivery to M cells or PP tissue. Proteins comprising chimerics of calreticulin plus a polypeptide with an antigen of choice would therefore prove valuable in that regard.

Members of the 14-3-3 protein family have been identified as regulatory elements in intracellular signaling pathways and cell cycle control. There had been reports that 14-3-3 protein can be used as a marker for Creutzfeldt-Jacob Disease (CJD) in cerebrospinal fluid (CSF). It is proposed that this protein or the gene coding for it is valuable in the control of the M cell phenotype, and as a result it would be advantageous to co-deliver that protein or gene with a protein, antigen, or DNA vaccine.

Nucleoside diphosphate kinases (NDP kinases) form a family of oligomeric enzymes present in all organisms. Eukaryotic NDP kinases are hexamers composed of identical subunits (approximately 17 kDa). A distinctive property of human NDPK-B is its ability to stimulate gene transcription. This property is independent of its catalytic activity and is possibly related to the role of this protein in cellular events including differentiation and tumor metastasis. Given our discovery of the increased expression of nucleoside diphosphate kinase B in M cell enriched PP FAE cells, we propose the importance of this protein in determining or controlling M cell phenotype, in M cell development, and optimal activation or priming of the mucosal immune system.

Example 5

Gene expression analysis of rat PP and NPP tissue samples

In addition to the proteomic studies highlighted above, PP and NPP tissue samples were sent for gene expression analysis to CLONTECH Laboratories Inc. (a division of Becton Dickinson (BD) Biosciences) who then extracted RNA from the tissues to probe on ATLASTM1.2 rat arrays. The data containing differential expression levels of 1,200 genes many of which are presented in Table 1 below. The data show over-expression of many proteins. In Table 1, over-expressed genes are shown in bold and italicized.

In Table 1, "N/C" means not calculated due to manually-determined inconsistencies in one or both spots, and "?" means low confidence level (small difference).

Also, over-expressed genes from Table 1 that had a fold change above 0, as well as over-expressed genes are shown in Table 2 below with corresponding GenBank accession numbers for rat and human origin.

Based on the results (ratio PP/ Normal epithelial tissue) in Table 1, the following proteins are of the particular interest: clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, Ca²⁺-dependant phospholipase A2 precursor, ribosomal proteins S12, S11, L12, L11, S29, S19, L21, L19, L13, L44, and L36A.

In addition a series of genes coding for different TFs was noted including the following:

Jun-B; c-jun related TF,

Jun-D; c-jun related TF,

STAT 3 - signal transducer and activator of transcription 3,

NF-kappa β Tf p105 subunit,

5 CREB active TF,
New england deaconess TF,
C-jun proto-oncogene; TF AP-1; RJG-9,
S-myc proto-oncogene; myc related,
Nm23-M2; nucleoside diphosphate kinase B; metastasis reducing protein,
NDK-B; nucleoside diphosphate kinase B ; metastasis reducing protein,
Lim-2; embryonic motor neuron topographic organizer; homeobox protein LIM-2, and
C-est-I proto-oncogene; p54.

10 TF coding genes such as these are considered here to be important in the
development of M cell phenotype and in priming the immune system. Their co-delivery or co-
targeting with DNA vaccine genes and/or with vaccines is expected to enhance activation of
mucosal immunity to the co-delivered DNA vaccine and/or antigen by virtue of their priming
of the cells to give a better mucosal immunity outcome.

15

TABLE 1							GENE EXPRESSION DATA FROM ATLAS 1.2 RAT ARRAY ANALYSIS	
#	coordinate	Spot Intensity		Ratio		Difference	GENE	
		PP1	NE1					
1	A03c	38	6	0.16	-32		T-cell surface glycoprotein CD5 precursor; lymphocyte glycoprotein LY1	
2	A03q	31	57	1.84	26		CD4 homologues, W3/25 antigen	
3	A04f	20	72	3.60	52		signal transducer CD24 precursor; heat stable antigen (HSA); nectadrin	
4	A04i	17	40	2.35	23		CD2, membrane glycoprotein, T-cell marker	
5	A04j	8	34	4.25	26		scavenger receptor class B type I	
6	A04m	16	31	1.94	15		SR13 myelin protein; peripheral myelin protein 22 (PMP-22); CD25 protein	
7	A05a	4	17	4.25	13		glutamyl aminopeptidase A	
8	A05i	52	29	0.56	-23		I-kB (I-kappa B) alpha chain; RLIF-1 gene product	
9	A05m	6	18	3.00	12		interferon regulatory factor 1 (IRF1)	
10	A06a	7	21	3.00	14		LIM domain protein CLP36, homologous to rat RIL	
11	A06c	152	78	0.51	-74		Gax, growth-arrest-specific protein	
12	A07c	42	24	0.57	-18		G1/S-specific cyclin D3 (CCND3)	
13	A07i	17	3	0.18	-14		M-phase inducer phosphatase 2 (MPI2); cell division control protein 25 B (CDC25B)	
14	A07n	37	20	0.54	-17		p55cdc; cell division control protein 20	
15	A08e	56	24	0.43	-32		prothymosin-alpha (PTMA)	
16	A08n	32	105	3.28	73		antigen peptide transporter 1	
17	A09q	13	36	2.77	23		proteasome delta subunit precursor; macropain delta; multicatalytic endopeptidase complex delta;	
18	A09i	15	27	1.80	12		proteasome component C13 precursor; macropain subunit C13; multicatalytic endopeptidase	
19	A09j	341	893	2.62	552		apolipoprotein A-I precursor (APO-AI)	
20	A08k	72	723	10.04	651		apolipoprotein A-IV precursor (APO-AIV)	
21	A11i	1	13	13.00	12		ErbB3 EGF receptor-related proto-oncogene; HER3	
22	A12c	21	48	2.29	27		A-raf proto-oncogene	
23	A12i	19	45	2.37	26		rac-alpha serine/threonine kinase (RAC-PK-alpha); protein kinase B (PKB); AKT1	
24	A13k	208	60	0.29	-148		HSP84; HSP90-beta; heat shock 90kD protein	
25	A14c	7	38	5.43	31		glutathione S-transferase Ya subunit (GST YA); ligandin subunit 1 alpha	
26	A14d	65	162	2.49	97		microsomal glutathione S-transferase (GST12; MGST1)	
27	A14e	94	160	1.70	66		glutathione S-transferase Yb subunit; GST subunit 4 mu (GSTM2)	
28	A14g	129	265	2.05	136		glutathione S-transferase P subunit; GST subunit 7 pi (GST7-7)	
29	A14n	7	19	2.71	12		NADPH-cytochrome P450 reductase (CPR); POR	
30	B01h	121	207	1.71	88		copper-zinc-containing superoxide dismutase 1 (Cu-Zn SOD1)	
31	B02f	4	25	6.25	21		fructose (glucose) transporter	
32	B05k	13	36	2.77	23		sodium channel SCN2B, beta 2 subunit, brain	
33	B06d	28	12	0.43	-18		potassium channel, inward rectifier 11	
34	B07k	2	18	9.00	16		proton-coupled dipeptide cotransporter	
35	B08b	59	181	3.07	122		fibroblast ADP/ATP carrier protein; ADP/ATP translocase 2; adenine nucleotide translocator 2	
36	B09i	7	43	6.14	36		sodium-glucose cotransporter 1	
37	B09l	40	115	2.88	75		Na+/K+ ATPase alpha 1 subunit	
38	B09n	43	78	1.81	35		vacuolar ATP synthase 16-kDa proteolipid subunit; ATP6C; MVP; ATP6	
39	B10b	10	78	7.80	68		sodium/potassium-transporting ATPase beta 1 subunit (ATP1B1)	
40	B12e	10	68	6.80	58		urate transporter/channel	
41	B12f	143	297	2.08	154		ATP synthase lipid-binding protein P1 precursor; ATPase protein 9; ATP5G1	
42	B12g	21	112	5.33	91		ATP synthase, subunit c, P2 gene	
43	B12m	18	38	2.11	20		annexin IV(ANX4); lipocortin IV-36-kDa zymogen granule membrane-associated protein (ZAP36)	
44	B13f	37	127	3.43	90		lipocortin 2	
45	B14d	59	326	5.53	267		fatty acid-binding protein (liver, L-FABP); Z-protein; squalene- & sterol-carrier protein (SCP); P14	
46	B14f	49	157	3.20	108		fatty acid-binding protein (intestinal, I-FABP; FABPI)	

#	coordinate	Spot Intensity		RATIO	Difference	GENE
		PP1	NE1			
47	C02f	19	78	4.11	59	fructose-bisphosphate aldolase B (ALDOB); liver-type aldolase
48	C02g	37	76	2.05	39	fructose-bisphosphate aldolase A (ALDOA); muscle-type aldolase
49	C02j	15	30	2.00	15	testis fructose-6-phosphate 2-kinase/fructose 2,6-bisphosphatase (PFKFB2); 6-phosphofructose-6-phosphate 2-kinase/fructose 2,6-bisphosphatase (PFKFB2)
50	C02h	804	1123	1.86	519	cytochrome c oxidase subunit Vb & V1a precursor (COX5B)
51	C03a	9	20	2.22	11	cytochrome B5 (CYB5)
52	C03e	65	138	2.12	73	mitochondrial hydroxymethylglutaryl-CoA synthase precursor (HMG-CoA synthase); 3-hydroxy-3-methylglutaryl-CoA synthase precursor (HMG-CoA synthase)
53	C03f	177	760	4.29	583	cytochrome oxidase, subunit I, Sertoli cells
54	C03g	27	47	1.74	20	ATPase, subunit F, vacuolar (vati)
55	C03i	641	1087	1.70	446	cytochrome c oxidase, subunit IV, mitochondrial
56	C03j	46	147	3.20	101	cytochrome c oxidase, subunit Va, mitochondrial
57	C03l	317	180	0.57	-137	cytochrome c oxidase, subunit VIII
58	C04b	189	351	1.86	162	mitochondrial ATP synthase beta subunit precursor (ATP5B)
59	C04g	23	127	5.52	104	creatine kinase, ubiquitous, mitochondrial
60	C08c	6	52	8.67	46	fatty acid amide hydrolase
61	C06l	73	175	2.40	102	cytochrome P450 17 (CYP17); P450C17; CYPXVII; steroid 17-alpha-hydroxylase/17,20 lyase
62	C07i	9	51	5.67	42	cytochrome P-450 4F1, hepatic tumour
63	C07j	19	39	2.05	20	cytochrome P-450 4F4
64	C08a	17	33	1.94	16	cytochrome P-450 4F5
65	C08l	8	24	3.00	16	adenylate kinase 3
66	C08m	32	65	2.03	33	cAMP-dependent protein kinase type I-alpha regulatory chain
67	C09e	7	30	4.28	23	glutathione synthetase (GSH synthetase; GSH-S; GSS); glutathione synthase
68	C10e	25	10	0.40	-15	carbonic anhydrase 4
69	C10j	19	9	0.47	-10	alkaline phosphatase
70	C10k	21	5	0.24	-18	dopamine beta-hydroxylase
71	C10l	29	11	0.38	-18	acetylcholinesterase, T subunit, glycolipid-anchored
72	C10m	228	445	1.95	217	NADPH+ alcohol dehydrogenase; aldehyde reductase (ALR); 3-dG-reducing enzyme
73	C11b	19	33	1.74	14	calcium binding protein 2 (CABP2); endoplasmic reticulum stress protein (ERP72); protein disulfide isomerase
74	C11d	97	55	0.57	-42	60S ribosomal protein L44; L36A
75	C11e	1728	527	0.30	-1201	40S ribosomal protein S12
76	C11g	400	179	0.45	-221	ribosomal protein L11
77	C11h	670	355	0.53	-315	ribosomal protein L13
78	C11i	867	424	0.44	-543	ribosomal protein L12
79	C11k	812	413	0.51	-399	S19; 40S ribosomal protein S19
80	C11l	554	286	0.52	-268	60S ribosomal protein L21
81	C11m	1405	737	0.52	-668	60S ribosomal protein L19 (RPL19)
82	C11n	315	109	0.35	-206	40S ribosomal protein S11
83	C12b	158	48	0.30	-110	Fte-1; putative v-fos transformation effector protein; yeast mitochondrial protein import homolog; 40S ribosomal protein S11
84	C12d	1451	787	0.54	-664	elongation factor 2 (EF2)
85	C12l	346	10	0.03	-336	clusterin (CLU); testosterone-repressed prostate message 2 (TRPM2); apolipoprotein J; sulfated
86	C12m	14	0	Undefined	-14	activator of apoptosis harakiri (HAKI); neuronal death protein 5 (DP5); BID3
87	C13c	19	5	0.26	-14	SURVIVAL OF MOTOR NEURON (SRM)
88	D08l	6	17	2.83	11	retinoid X receptor alpha (RXR alpha; RXRA); NR2B1
89	D12c	6	18	3.00	12	INOSITOL TRIPHOSPHATE RECEPTOR SUBTYPE 3
90	E01m	10	50	5.00	40	neurotrophin 3 precursor (NTF3); neurotrophic factor; HDNF; nerve growth factor 2 (NGF2)
91	E02c	19	8	0.42	-11	transforming growth factor, beta 1
92	E03k	40	185	4.63	145	C-type natriuretic peptide precursor (CNP; NPPC)
93	E04b	17	7	0.41	-10	thyroid stimulating hormone, beta
94	E07c	33	15	0.45	-18	c-src-kinase (CSK) & negative regulator; tyrosine-protein kinase

#	coordinate	Spot Intensity		RATIO	Difference	GENE
		PP1	NE1			
95	E08a	105	249	2.37	144	extracellular signal-regulated kinase 1 (ERK1); mitogen-activated protein kinase 1 (MAP kinase 1; protein kinase C delta type (PKC-delta)
96	E08i	8	22	2.75	14	insulin receptor-related receptor-alpha (sIRR-1)
97	E09j	36	68	1.89	32	CamK II; calcium/calmodulin-dependent protein kinase brain type II beta
98	E10f	3	17	5.67	14	CamK I; calcium/calmodulin-dependent protein kinase type I + CaM-like protein kinase
99	E10q	3	13	4.33	10	Casein kinase I delta; CKId; 49-kDa isoform
100	E10j	18	42	2.33	24	cyclin-dependent kinase 4 (CDK4); cell division protein kinase 4; PSK-J3
101	E11e	15	29	1.93	14	protein phosphatase 2C isoform; Mg2+ dependent protein phosphatase beta isoform
102	E12q	23	45	1.96	22	<i>guanine nucleotide-binding protein G(i) alpha 2 subunit (GNAI2); adenylate cyclase-</i>
103	E13i	40	19	0.48	-21	rab12, ras related GTPase
104	E14i	268	661	2.47	393	RalGDSB; GTP/GDP dissociation stimulator for a ras-related GTPase
105	F01e	10	29	2.90	19	<i>calcium-dependent phospholipase A2 precursor (PLA2); phosphatidylcholine 2-</i>
106	F02a	189	97	0.51	-92	phospholipase C beta 3 (PLC-beta 3)
107	F02h	15	79	5.27	64	14-3-3 protein zeta/delta; PKC inhibitor protein-1; KCIP-1; mitochondrial import stimulation factor
108	F04e	33	57	1.73	24	14-3-3 protein epsilon; PKC inhibitor protein-1; KCIP-1; mitochondrial import stimulation factor L
109	F04k	15	29	1.93	14	presenilin 1 (PSNL1; PSEN1; PS1); S182 protein
110	F05e	8	21	2.63	13	PDGF-associated protein
111	F05i	6	17	2.83	11	ADP-ribosylation factor 5 (ARF5)
112	F06a	59	143	2.42	84	dipeptidase (DPEP1)
113	F06j	12	22	1.83	10	granzyme M precursor (GZMM); MET-ASE; natural killer cell granular protease; RNK-MET-1
114	F07m	7	29	4.14	22	angiotensin converting enzyme (ACE; somatic; dipeptidyl carboxypeptidase I; kininase II
115	F08e	11	100	9.09	89	amorphinase B
116	F08h	18	48	2.67	30	kidney aminopeptidase M (APM)
117	F08j	39	92	2.36	53	metalloendopeptidase meprin beta subunit
118	F08k	19	81	4.26	62	endothelin converting enzyme
119	F08l	5	15	3.00	10	gelatinase A
120	F09a	3	16	5.33	13	cathepsin L
121	F09q	13	27	2.08	14	proteasome component C3
122	F09l	14	28	2.00	14	leukocyte common antigen-related tyrosine phosphatase (LAR)
123	F12a	6	16	2.67	10	ornithine decarboxylase (ODC)
124	G31	12	39	3.25	27	cytoplasmic beta-actin (ACTB)
125	G43	287	1132	3.94	845	<i>40S ribosomal protein S29 (RPS29)</i>
126	G47	7327	3614	0.49	-3713	

Table 2
RAT GENES (PP VS. NPP)

GENE	Fold change	GenBank ID Human	GenBank ID Rat
activator of apoptosis harakiri (HRK); neuronal death protein 5 (DP5); BID3	**	U76376.1	D83697
RET ligand 1 (RET1)	**	-	U97142
P2X purinoceptor 1; ATP receptor P2X1; purinergic receptor; RP-2 protein	**	P51575	U14414
leukocyte common antigen precursor (LCA); CD45 antigen; T200; PTPRC	**	Y00638	M10072
amphiphysin II (AMPH2)	**	AF001383.1	Y13380
Jak3 tyrosine-protein kinase; Janus kinase 3	**	XM_038595.3	D28508
DCC; netrin receptor; immunoglobulin gene superfamily member; former tumor suppressor protein candidate	**	M32292.1	AH002168.1
c-fgr proto-oncogene	**	AAA52762.1	X57018.1
small inducible cytokine A3 precursor (SCYA3); macrophage inflammatory protein 1 alpha precursor (MIP1-alpha; MIP1A)	**	P10147	AF119381.1
protein kinase C beta-I type (PKC-beta I) + protein kinase C beta-II type (PKC-beta II)	**	X06318	P04410
E-selectin precursor; endothelial leukocyte adhesion molecule 1 (ELAM-1); leukocyte-endothelial cell adhesion molecule 2 (LECAM2); CD62E	**	P16581	L25527
T-cell receptor CD3 zeta subunit	**	J04132.1	L08447.1
Protein kinase C-binding protein beta15; RING-domain containing	**	-	U48248
G1/S-specific cyclin C (CCNC)	**	AAC50825.1	D14013
maspin; protease inhibitor 5 (PI5); tumor suppressor	**	U04313.1	U58857
peptide/histidine transporter	**	-	AB000280
acetyl-CoA carboxylase (ACC); biotin carboxylase	**	X68968.1	AH002123.1
fibroblast growth factor receptor subtype 4	**	L03840.1	M91599
LCR-1; putative chemokine and HIV coreceptor homolog; G protein-coupled receptor	**	-	U54791
tumor necrosis factor alpha precursor (TNF-alpha; TNFA); cachectin	**	AF043342.1	X66539
CC chemokine MIP3 alpha exodus	**	-	U90447.1
lutetizing hormone, alpha	**	NM_000735.2	V01252
Ctk; non-receptor protein tyrosine kinase (batk)	**	P42679	L34542.1
RhoGAP; p122	**	-	S54293
Adenylyl cyclase type V	**	M83533.1	M96159
cathepsin S precursor (CTSS)	**	P25774	L03201.1
O-6-methylguanine-DNA methyltransferase (MGMT); methylated-DNA-protein-cysteine methyltransferase	**	M31767.1	NM_012861.1
clusterin (CLU); testosterone-repressed prostate message 2 (TRPM2); apolipoprotein J; sulfated glycoprotein 2 (SGP2); dimeric acid glycoprotein (DAG)	34.60	X14723	U02391.1
T-cell surface glycoprotein CD5 precursor; lymphocyte glycoprotein LY-1 (LYT1)	6.33	X04391.1	D10728
M-phase inducer phosphatase 2 (MPI2); cell division control protein 25 B (CDC25B)	5.67	S78187.1	D16237
dopamine beta-hydroxylase	4.20	Y00096.1	L12407

GENE	Fold change	GenBank ID Human	GenBank ID Rat
SURVIVAL OF MOTOR NEURON(RSMN)	3.80	AAC50473.1	U75369
HSP84; HSP90-beta; heat shock 90kD protein	3.47	XM_055551.3	S45392
Fte-1; putative v-fos transformation effector protein; yeast mitochondrial protein import homolog; 40S ribosomal protein S3A ; RPS3A	3.29	M84711.1	M84716.1
40S ribosomal protein S12	3.28	X53505	M18547
40S ribosomal protein S11	2.89	X06617	K03250
acetylcholinesterase, T subunit, glycolipid-anchored	2.64	M55040.1	X71089.1
carbonic anhydrase 4	2.50	NM_000717.2	I52551
thyroid stimulating hormone, beta	2.43	S70587.1	M13897.1
transforming growth factor, beta 1	2.38	M34057	NM_021578.1
prothymosin-alpha (PTMA)	2.33	AF257099.1	M20035
potassium channel, inward rectifier 11	2.33	-	D42145
ribosomal protein L12	2.28	L06505.1	X53504.1
ribosomal protein L11	2.23	X79234.1	X62146.1
c-src-kinase (CSK) & negative regulator; tyrosine-protein kinase	2.20	X59932.1	X58631
alkaline phosphatase	2.11	AAA98616.1	S18408
guanine nucleotide-binding protein G(i) alpha 2 subunit (GNAI2); adenylate cyclase-inhibiting G alpha protein	2.11	XM_041507.1	NM_031035.1
40S ribosomal protein S29 (RPS29)	2.03	NM_001032.2	X59051
S19; 40S ribosomal protein S19	1.97	P39019	P17074
Gax, growth-arrest-specific protein	1.95	-	Z17223.1
calcium-dependent phospholipase A2 precursor (PLA2); phosphatidylcholine 2-acylhydrolase (PLA2-10; PLA2G5)	1.95	M22430.1	U38376.1
60S ribosomal protein L21	1.94	P46778	M27905
60S ribosomal protein L19 (RPL19)	1.91	X63527	J02650
ribosomal protein L13	1.89	P26373	X78327.1
p55cdc; cell division control protein 20	1.85	AF099644.1	AF052695.1
elongation factor 2 (EF2)	1.84	X51466	Y07504.1
I-kB (I-kappa B) alpha chain; RL/IF-1 gene product	1.79	X63594.1	AF388201.1
60S ribosomal protein L44; L36A	1.76	M15661	P10661
cytochrome c oxidase, subunit VIIIh	1.76	J04823.1	NM_012786.1
G1/S-specific cyclin D3 (CCND3)	1.75	NM_001760.2	NM_012766.1
cytochrome c oxidase, subunit IV, mitochondrial	0.59	AF017115.1	X14209
glutathione S-transferase Yb subunit; GST subunit 4 mu (GSTM2)	0.59	-	X04229.1
copper-zinc-containing superoxide dismutase 1 (Cu-Zn SOD1)	0.58	-	NM_017050.1
14-3-3 protein zeta/delta; PKC inhibitor protein-1; KCIP-1; mitochondrial import stimulation factor S1 subunit	0.58	U28964.1	L07913.1
calcium binding protein 2 (CABP2); endoplasmic reticulum stress protein (ERP72); protein disulfide isomerase-related protein precursor	0.58	XM_012077.4	M86870
ATPase, subunit F, vacuolar (vaf)	0.57	AF047436.1	U43175.1
proteasome component C13 precursor; macropain subunit C13; multicatalytic endopeptidase complex subunit C13; PSMB8	0.56	P28062	NM_080767.1
vacuolar ATP synthase 16-kDa proteolipid subunit; ATP6C; MVP; ATPL	0.55	NM_001695.1	M62762.1

GENE	Fold change	GenBank ID Human	GenBank ID Rat
dipeptidase (DPEP1)	0.55	NM_004413.1	M94056
CD4 homologue, W3/25 antigen	0.54	BC025782.	M15768.1
mitochondrial ATP synthase beta subunit precursor (ATP5B)	0.54	NM_001686.1	M19044.1
cytochrome c oxidase subunit Vb & VIa precursor (COX5B)	0.54	M59250.1	X14208.1
insulin receptor-related receptor-alpha (sIRR-1)	0.53	-	M90660.1
cyclin-dependent kinase 4 (CDK4); cell division protein kinase 4; PSK-J3	0.52	P11802	P35426
14-3-3 protein epsilon; PKC inhibitor protein-1; KCIP-1; mitochondrial import stimulation factor L subunit	0.52	XM_088041.1	D30739.1
SR13 myelin protein; peripheral myelin protein 22 (PMP-22); CD25 protein	0.52	-	M69139.1
cytochrome P-450 4F5	0.52	-	AF288818.1
NADP+ alcohol dehydrogenase; aldehyde reductase (ALR); 3-dG-reducing enzyme	0.51	J04794.1	D10854.1
protein phosphatase 2C isoform; Mg2+ dependent protein phosphatase beta isoform	0.51	-	S90449.1
testis fructose-6-phosphate 2-kinase/fructose 2,6-biphosphate (testis 6PF-2-K/fru-2,6-P2ase); 6-phosphofructo- 2-kinase; fructose-2,6-bisphosphatase	0.50	NM_002625.1	X15579.1
proteasome component C3	0.50	D00760	J02897.1
cAMP-dependent protein kinase type I-alpha regulatory chain	0.49	P10644	P09456
cytochrome P-450 4F4	0.49	-	U39206.1
fructose-bisphosphate aldolase A (ALDOA); muscle-type aldolase	0.49	XM_043948.2	NM_012495.1
glutathione S-transferase P subunit; GST subunit 7 pi (GST7-7)	0.49	-	X02904.1
ATP synthase lipid-binding protein P1 precursor; ATPase protein 9; ATP5G1	0.48	NM_005175.1	NM_017311.1
cathepsin L	0.48	M20496.1	Y00697.1
annexin IV(ANX4); lipocortin IV;36-kDa zymogen granule membrane-associated protein (ZAP36)	0.47	XM_031596.3	NM_024155.1
mitochondrial hydroxymethylglutaryl-CoA synthase precursor (HMG-CoA synthase); 3-hydroxy-3-methylglutaryl-CoA synthase; HMGCS2	0.47	P54868	P22791
cytochrome B5 (CYB5)	0.45	M22865.1	D13205.1
A-raf proto-oncogene	0.44	P10398	X06942
Casein kinase I delta; CKId; 49-kDa isoform	0.43	P48730	Q06486
CD2, membrane glycoprotein, T-cell marker	0.43	M14362.1	X05111.1
kidney aminopeptidase M (APM)	0.42	XM_087746.1	M26710
rac-alpha serine/threonine kinase (RAC-PK-alpha); protein kinase B (PKB); AKT1	0.42	P31749	Y15748.1
extracellular signal-regulated kinase 1 (ERK1); mitogen-activated protein kinase 1 (MAP kinase 1; MAPK1); insulin- stimulated microtubule-associated protein-2 kinase; MNK1; PRKM3; ERT2; p44-MAPK	0.42	P27361	P21708
cytochrome P450 17 (CYP17); P450C17; CYPXVII; steroid 17-alpha-hydroxylase/17,20 lyase	0.42	NM_000102.2	X69816.1
ADP-ribosylation factor 5 (ARF5)	0.41	NM_001662.	NM_024149.1

GENE	Fold change	GenBank ID Human	GenBank ID Rat
rab12, ras related GTPase	0.41	-	M83676.
microsomal glutathione S-transferase (GST12; MGST1)	0.40	XM_048886.3	J03752
apolipoprotein A-I precursor (APO-AI)	0.38	X02162	M00001
presenilin 1 (PSNL1; PSEN1; PS1); S182 protein	0.38	XM_007441.1	D82363
amonipeptidase B	0.38	XM_087242.1	U61696
leukocyte common antigen-related tyrosine phosphatase (LAR)	0.38	-	U00477.1
NADPH-cytochrome P450 reductase (CPR); POR	0.37	S90469	NM_031576.1
protein kinase C delta type (PKC-delta)	0.36	NM_006254.1	M18330
proteasome delta subunit precursor; macropain delta; multicatalytic endopeptidase complex delta; proteasome subunit Y; proteasome subunit 5; PSMB6	0.36	X61971.1	NM_057099.1
sodium channel SCN2, beta 2 subunit, brain	0.36	AAC05208.1	NM_012877.
retinoid X receptor alpha (RXR alpha; RXRA); NR2B1	0.35	XM_088424.1	NM_012805.1
PDGF-associated protein	0.35	U41745.1	U41744.1
Na+/K+ ATPase alpha 1 subunit	0.35	AAA51801.1	M28647
RaGDSB; GTP/GDP dissociation stimulator for a ras-related GTPase	0.34	-	NM_019250.1
interferon regulatory factor 1 (IRF1)	0.33	XM_034862.1	M34253
LIM domain protein CLP36, homologous to rat RIL	0.33	AJ310549.1	U23769.1
adenylate kinase 3	0.33	XM_016642.3	NM_013218.1
INOSITOL TRIPHOSPHATE RECEPTOR SUBTYPE 3	0.33	-	L06096.1
endothelin converting enzyme	0.33	Z35307.1	D29683
fibroblast ADP/ATP carrier protein; ADP/ATP translocase 2; adenine nucleotide translocator 2 (ANT2)	0.33	J02683	D12771
cytochrome c oxidase, subunit Va, mitochondrial	0.31	M22760.1	X15030
fatty acid-binding protein (intestinal; I-FABP; FABPI)	0.31	M18079	M18080.1
ornithine decarboxylase (ODC)	0.31	X16277	D11372.1
antigen peptide transporter 1	0.30	X57522	P36370
lipocortin 2	0.29	D00017.1	S73557
signal transducer CD24 precursor; heat stable antigen (HSA); nectadrin	0.28	P25063	U49062
cytoplasmic beta-actin (ACTB)	0.25	M10277.1	V01217
fructose-bisphosphate aldolase B (ALDOB); liver-type aldolase	0.24	XM_042788.1	M10149
granzyme M precursor (GZMM); MET-ASE; natural killer cell granular protease; RNK-MET-1	0.24	NM_005317.2	Q03238
scavenger receptor class B type I	0.24	-	AB002151.1
glutamyl aminopeptidase A	0.24	XM_003595.2	S73583
metalloendopeptidase meprin beta subunit	0.23	-	M88601.1
glutathione synthetase (GSH synthetase; GSH-S; GSS); glutathione synthase	0.23	U34683.1	L38615.1
cytochrome oxidase, subunit I, Sertoli cells	0.23	S79304	S79304
CamK I; calcium/calmodulin-dependent protein kinase type I + CaM-like protein kinase	0.23	Q14012	L24907
C-type natriuretic peptide precursor (CNP; NPPC)	0.22	NM_024409.1	D90219

GENE	Fold change	GenBank ID Human	GenBank ID Rat
neurotrophin 3 precursor (NTF3); neurotrophic factor; HDNF; nerve growth factor 2 (NGF2)	0.20	M37763.1	M34643
phospholipase C beta 3 (PLC-beta 3)	0.19	NM_000932.1	M99567
ATP synthase, subunit c, P2 gene	0.19	D13119.1	D13124
gelatinase A	0.19	NM_004530.1	U65656
glutathione S-transferase Ya subunit (GST YA); ligandin subunit 1 alpha	0.18	NM_000852.2	K01932
creatine kinase, ubiquitous, mitochondrial	0.18	XM_016524.4	X59737
fatty acid-binding protein (liver; L-FABP); Z-protein; squalene- & sterol-carrier protein (SCP); P14	0.18	NM_001443.1	M35991
cytochrome P-450 4F1, hepatic tumour	0.18	-	NM_019623.1
CamK II; calcium/calmodulin-dependent protein kinase brain type II beta	0.18	NM_001220.1	M16112
sodium-glucose cotransporter 1	0.16	P13866	U03120
fructose (glucose) transporter	0.16	AAB60641	D13871.1
urate transporter/channel	0.15	-	U67958
sodium/potassium-transporting ATPase beta 1 subunit (ATP1B1)	0.13	NM_001677.1	NM_013113.1
fatty acid amide hydrolase	0.12	U82535.1	U72497
proton-coupled dipeptide cotransporter	0.11	-	D50306.1
angiotensin converting enzyme (ACE; somatic; dipeptidyl carboxypeptidase I; kininase II	0.11	NM_000789.1	NM_012544.1
apolipoprotein A-IV precursor (APO-AIV)	0.10	XM_052144.2	P02651
ErbB3 EGF receptor-related proto-oncogene; HER3	0.08	M29366.1	NM_017218.2
Jun-B; c-jun related TF,	**	M29039.1	X54686
S-myc proto-oncogene; myc related,	**		M29069
C-est-I proto-oncogene; p54.	5	AF193068.1	X55787.1
Jun-D; c-jun related TF,	1.79	X56681.1	D26307(mouse)
NF-kappa β Tf p105 subunit,	1.67	P19838	L26267.1
Nm23-M2; nucleoside diphosphate kinase B; metastasis reducing protein,	1.47		X68193.1 (mouse)
STAT 3 - signal transducer and activator of transcription 3,	1.16	NM_003150.1	NM_012747.1
CREB active TF,	1		M34356.1
New england deaconess TF,	1		U09229
Lim-2; embryonic motor neuron topographic organizer; homeobox protein LIM-2, and	1		L35572
NDK-B; nucleoside diphosphate kinase B ; metastasis reducing protein,	0.81		U29200.1
C-jun proto-oncogene; TF AP-1; RJG-9,	0.5	J04111.1	X17215.1

Symbols indicating fold changes in Table 2:

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- ** : expressed in PP but not NPP, or in co-culture but not in Caco-2 cells.
 * : expressed in co-culture but not in Caco-2 cells (only repeated once).
 - : expressed in Caco-2 but not in co-culture.

Example 5ATLAS array data on co-culture of human Caco-2 cells and Raji B-cells

5 In order to facilitate the routine study of M cell biology, there was a desire to establish
a suitable and representative in-vitro model. In the work carried out by Kernéis *et al.*
(Kerneis S, Caliot E, Stubbe H, Bogdanova A, Kraehenbuhl J, Pringault E (2000). Molecular
studies of the intestinal mucosal barrier physiopathology using co-cultures of epithelial and
immune cells: a technical update. *Microbes Infect* 2000 Jul;2 (9):1119-24), it was reported
10 that Peyer's patch lymphocytes co-cultured with Caco-2 cells trigger the phenotypic
conversion of enterocytes into cells that express morphological and functional M-cell
properties. This work was further developed by Gullberg *et al.* (Gullberg E, Leonard M,
Karlsson J, Hopkins AM, Brayden D, Baird AW, Artursson P. Expression of specific markers
and particle transport in a new human intestinal M-cell model. *Biochem Biophys Res*
15 *Commun* 2000 Dec 29; 279(3): 808-13) to create a simplified in vitro model of the human M-
cell. Co-cultures of physically separated human intestinal epithelial Caco-2 cells and B-cell
lymphoma Raji cells were established. The co-cultures were characterized under the criteria
of morphology, integrity, expression of M-cell markers and cell adhesion molecules (CAMs),
and altered particle transport. Using this construct, the epithelial cells were transformed to
20 cells with an M-cell-like morphology and had altered expression of potential human M-cell
markers (alkaline phosphatase down-regulation and Sialyl Lewis A antigen up-regulation).
The expression of intercellular adhesion molecule-1 and vascular cell adhesion molecule
was altered, and there was an increased binding of lectins wheat germ agglutinin and peanut
agglutinin with a 40-fold increase in microparticle transport. The particle transport was size-
25 dependent and could be inhibited at 4°C or by replacing the Raji B-cells with Jurkat T-cells.
Thus the comparison of RNA isolated from co-cultured Caco2 cells to that isolated from
normal Caco2 cells was designed to simulate a comparison of M cell RNA to normal gut
enterocyte RNA.

30 Isolation of total RNA from Co-Cultured Caco-2 cells

Caco-2 cell culture

Caco-2 cells were cultured in Dulbecco's Modified Eagles Medium (DMEM), 4.5g/L
glucose supplemented with 1% Mem, 10% FCS and 1% penicillin/streptomycin at 37°C and

5% CO₂ in 95% relative humidity. Cells were grown and expended in Falcon culture flasks and passaged once they attained 100% confluence. Caco-2 cells were seeded on Transwell Clear filters (Costar, 12mm diameter, 3.0um pore size)) at a density of 5x10⁵ cells/cm² and incubated in a 12 well culture plate with a medium change every second day. 1.0ml was added to the basolateral side and apical sides.

Raji cell culture

Raji B-lymphoma cells were cultured in RPMI 1640 Medium, with 1% (v/v) non-essential amino-acids, 10% FCS and 1% penicillin/streptomycin, 1% L-glutamine at 37°C and 5% CO₂ in 95% relative humidity. Cells were grown in suspension in Falcon tissue culture flasks and passaged by dilution every 5-7 days.

Co-culture: day 14 (treating with Raji B-cells)

After 14 days of culturing Caco-2 monolayers, 15-20ml of Raji cells were removed from the T75 flask and placed in a 20ml universal. The cells were centrifuged at 1000 rpm for 3 min. Cells were re-suspended at a concentration of 1x10⁶ cells/ml. 1ml of fresh complete DMEM was added to the apical and basolateral sides of the Caco-2 monolayer filters. 0.5 ml of 1x10⁶ Raji cells/ml cells was added to the basolateral side of the filters. For control filters (non co-culture) 0.5ml of Raji medium only was added to the basolateral side.

Isolation of Total RNA from co-cultured Caco-2 cells

After 4 days of co-culture the filters were rinsed in PBS. 0.5 ml of PBS was added to the apical side of each filter and the Caco-2 cells were scraped off the filter surface into suspension in the PBS. The cells from all the co-cultured Caco-2 filters were pooled, centrifuged at 1000 rpm for 3 min, the supernatant PBS was removed and the pellet was used for RNA extraction.

Analysis of mRNA expression

Total cellular RNA was extracted using an acid guanidinium thiocyanate-phenol-chloroform method. RNA's integrity was confirmed by gel electrophoresis and ethidium bromide staining. mRNA was reverse transcribed in the presence of P³² dATP, and the transcribed cDNA was purified by chromatography before being hybridized over night to the array membrane. Membranes were exposed to x-ray film using an intensifying screen for 3 days and the mRNA expression levels were analyzed by scanning the films with a

densitometer. Expression levels were normalized relative to internal standards, and relative increases in mRNA levels in co-cultured cells versus monoculture controls were calculated. Two hybridization experiments were performed using mRNA from two separate cell harvests. Results from the two experiments were pooled, and a summary of the findings was tabulated in Tables 3(a)-3(f). The identified genes are from the following groups: oncogenes, tumor suppressor genes, genes involved in the cell cycle, ion channels and transport, stress response genes, modulators and effectors, genes involved in intracellular transduction, genes linked to apoptosis, DNA synthesis, repair & recombination, transcription factors, DNA binding proteins, receptors, cell surface antigens, genes involved in cell adhesion, growth factors, cytokines, chemokines and hormones.

In Table 3, genes which were found to be exclusively over-expressed in the co-culture and not in the control Caco-2 monolayer are represented by **. A single asterisk represents genes that also were expressed in the co-culture and not in the control Caco-2 monolayer. However, these particular genes have been distinguished from the genes labeled with two asterisks as they were not expressed in both hybridization experiments performed, and will require confirmation in the future by PCR so as to rule out false positives/negatives. Genes not expressed in the co-culture but expressed in the Caco-2 monolayer controls are indicated by a minus symbol, "-".

Table 3 a: Oncogenes, Tumor Suppressors, Cell Cycle Regulators

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
Myeloid cell nuclear differentiation antigen (MNDA)	*	M81750
G1/S-specific cyclin D1 (CCND1); cyclin parathyroid adenomatosis 1 (PRAD1); bcl-1 oncogene	*	X59798
cyclin-dependent kinase 4 inhibitor 2 (CDK4I; CDKN2); p16-INK4; multiple tumor suppressor 1 (MTS1)	*	L27211.1
cyclin-dependent kinase inhibitor 1C (CDKN1C); p57-KIP2	*	U22398
ezrin; cytovillin 2; villin 2 (VIL2)	1.69	X51521
proto-oncogene tyrosine-protein kinase kit; c-kit; mast/stem cell growth factor receptor precursor(SCFR); CD117 antigen	1.55	L04143.1
proliferating cell nucleolar antigen P120; NOL1	1.52	M32110
jun proto-oncogene; avian sarcoma virus 17 oncogene homolog; transcription factor AP-1	1.47	J04111
C-src proto-oncogene (SRC1)	1.35	X59932
CDC-like kinase 3 (CLK3)	1.35	L29220
cell division cycle protein 25 nucleotide exchange factor (CDC25)	1.34	M91815.1
prothymosin alpha (PROT-alpha; PTMA)	1.32	M26708
40S ribosomal protein S19 (RPS19)	1.31	M81757
avian myelocytomatosis viral oncogene homolog (MYC)	1.30	V00568
CDC-like kinase 1 (CLK1)	1.27	L29219.1
cyclin-dependent kinase 4 inhibitor 2D (CDKN2D); p19-INK4D	0.69	U49399.1
vascular endothelial growth factor receptor 1 (VEGFR1); tyrosine-protein kinase receptor flt + soluble VEGFR; tyrosine-protein kinase receptor SFLT	0.62	XM_039993.2
neogenin	-	U61262.1
webB2 receptor protein-tyrosine kinase; neu proto-oncogene; c-erbB2 + HER2 receptor	-	M11730.1
N-ras; transforming p21 protein	-	AAA60255

Table 3 b: Ion Channels, Modulators, Effectors

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<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
extracellular signal-regulated kinase 3 (ERK3); mitogen-activated protein kinase 6 (MAP kinase 6; MAPK6; PRKM6); p97-MAPK	**	X14798.1
40-kDa heat-shock protein 1 (HSP40); DNAJ protein homolog 1 (HDJ1; DNAJ1)	**	D49547
70-kDa heat shock protein 1 (HSP70.1; HSPA1)	**	M11717
glutaredoxin	**	X76648
tyrosine kinase receptor tie-1 precursor	*	AAB84296
ras-related protein RAB3B	*	NM_002867.1
macMARCKS; MARCKS-related protein (MRP); MLP	*	P49006
mitogen-activated protein kinase 3 (MAPK3; PRKM3);	*	P27361

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
MAPK1; extracellular signal-regulated kinase 1 (ERK); microtubule-associated protein 2 kinase; insulin-stimulated MAP2kinase		
mitogen-activated protein kinase 9 (MAP kinase 9; MAPK9; PRKM9); c-jun N-terminal kinase 2 (JNK2); JNK55	*	NM_002752.1
60-kDa heat shock protein (HSP60); HSPD1; 60-kDa chaperonin; mitochondrial matrix protein P1 precursor; p60 lymphocyte protein; HUCHA60; GROEL	*	M22382.1
serine kinase	2.24	U09564.1
transferrin receptor (TFRC); CD71 antigen	1.80	M11507.1
Neurotrophic tyrosine kinase receptor-related 3: TKT precursor	1.63	U55017.1
phospholipase C (PLCL)	1.62	X14034.
cAMP-response element binding protein (CREB)	1.59	M27691.1
ephrin type-A receptor 1 precursor; tyrosine-protein kinase receptor eph	1.55	M18391
27-kDa heat-shock protein (HSP27); stress-responsive protein 27 (SRP27); estrogen-regulated 24-kDa protein; HSPB1	1.42	X54079.1
tyrosine kinase trk1	1.42	XM_012654.3
ras-related protein RAB3A	1.38	XM_054457.2
janus kinase 3 (JAK3); leukocyte janus kinase (L-JAK)	1.33	XM_038595.3
dual-specificity mitogen-activated protein kinase kinase 1 (MAP kinase kinase 1; MAPKK 1; MKK1); extracellular signal-regulated kinase 1; ERK activator kinase 1	1.29	NM_002755.2
calcium/calmodulin-dependent protein kinase type IV catalytic subunit (CAMK IV); CAM kinase-GR	1.27	NM_001744.1
ras-related protein RAB5A	0.75	XM_053461.2
colon carcinoma kinase 4 precursor (CCK4) + transmembrane receptor PTK7	0.68	U33635.1
epithelial discoidin domain receptor 1 precursor (EDDR1; DDR1); cell adhesion kinase (CAK); TRKE; RTK6; protein tyrosine kinase 3A (PTK3A); neuroepithelial tyrosine kinase (NEP)	0.63	XM_004559.5
ras-related protein RAB6	0.27	M28212.1
cAMP-dependent protein kinase type I beta regulatory subunit (PRKAR1B)	0.23	M65066.1
tyrosine-protein kinase ack	-	CAC15525
T-lymphocyte maturation-associated protein MAL	-	P21145
orphan hormone nuclear receptor	-	U04897.1
LIM domain kinase 1 (LIMK-1)	-	P53667
protein kinase C alpha polypeptide (PKC-alpha; PKCA)	-	NM_002737.1
dual specificity mitogen-activated protein kinase kinase 3 (MAP kinase kinase 3; MAPKK3; MKK3); ERK activator kinase 3; MAPK/ERK kinase 3 (MEK3)	-	P46734
Yamaguchi sarcoma viral-related oncogene homolog; tyrosine -protein kinase lyn	-	M16038.1
protein-tyrosine phosphatase 1E	-	U12128.1

Table 3.c: Apoptosis, DNA Synthesis, Repair & Recombination

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
ubiquitin-conjugating enzyme E2 17-kDa (UBE2A); ubiquitin-protein ligase; ubiquitin carrier protein, HR6A	**	NM_003336.1
growth arrest & DNA-damage-inducible protein 153 (GADD153); DNA-damage-inducible transcript 3 (DDIT3); C/EBP homologous protein (CHOP)	**	S40706.1
growth factor receptor-bound protein 2 (GRB2); ASH protein	*	M96995.1
glutathione S-transferase A1 (GTH1; GSTA1); HA subunit 1; GST-epsilon	*	M21758.1
cytoplasmic dynein light chain 1 (HDLC1); protein inhibitor of neuronal nitric oxide synthase (PIN)	*	U32944.1
xeroderma pigmentosum group G complementing protein (XPG); X-ray repair-complementing defective repair in Chinese hamster cells 5 (XRCC5)	*	NM_021141.2
xeroderma pigmentosum group D complementing protein (XPD); X-ray repair-complementing defective repair in Chinese hamster cells 2 (XRCC2)	*	AF035587.1
RAD23 homolog A (RAD23A; hHR23A)	*	NM_005053.1
ataxia telangiectasia (ATM)	*	AAB38309
apoptosis regulator bcl-x	1.60	Z23115.1
caspase 9 precursor (CASP9); ICE-like apoptotic protease 6 (ICE-LAP6); apoptotic protease MCH6; apoptotic protease activating factor 3 (APAF3)	1.42	AB020979.1
CD40 receptor-associated factor 1 (CRAF1)	1.39	U21092.1
SL cytokine precursor; FMS-related tyrosine kinase 3 ligand (FLT3 ligand; FLT3LG)	1.35	NM_001459.1
cytochrome P450 reductase	1.33	AAD45961.1
X-ray repair complementing defective repair in Chinese hamster cells 1 (XRCC1)	1.25	M36089
Ku (p70/p80) subunit; ATP-dependent DNA helicase II 86-kDa subunit; lupus ku autoantigen protein; thyroid-lupus autoantigen (TLAA); CTC box binding factor 85-kDa subunit (CTCBF; CTC85); nuclear factor IV	0.74	X57500.1
caspase 10 precursor (CASP10); ICE-LIKE apoptotic protease 4 (ICE-LAP4); apoptotic protease MCH4; fas-associated death domain protein; interleukin 1 beta-converting enzyme 2 (FLICE2);	0.45	Q92851
inhibitor of apoptosis protein 2 (IAP2; IAP2) + IAP homolog B; TNFR2-TRAF signaling complex protein 2; MIHB	-	Q13490
recA-like protein HsRad51; DNA repair protein RAD51 homolog	-	BAA02962.1
DNA damage repair & recombination protein 52 (RAD52)	-	B56529
DNA ligase III (LIG3); polydeoxyribonucleotide synthase	-	CAA59230.1

Table 3 d: Transcription Factors, DNA Binding Proteins

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
transcriptional activator hSNF2-alpha	**	D26155.1
early growth response protein 1 (EGR1); transcription factor ETR103; KROX24; zinc finger protein 225 (ZNF225); AT225	2.71	M62829.1
homeobox A1 protein (HOXA1); HOX1F	2.17	U10421.1
transcription factor NF-ATc	1.67	U08015.1
R kappa B DNA-binding protein	1.66	U08191.1
transcription initiation factor IID 31-kDa subunit (TFIID); TATA-box-binding protein-associated factor RNA polymerase II G 32-kDa subunit (TAFII32; TAF2G); TAFII31	1.57	M55654
homeobox protein hLim1; LHX1	1.51	NM_005568.1
helix-loop-helix protein HLH 1R21; DNA-binding protein inhibitor Id-3; HEIR-1	1.49	X69111.1
guanine nucleotide-binding protein G-s alpha subunit (GNAS); adenylate cyclase-stimulating G alpha protein	1.46	NP_000507.1
CCAAT-binding transcription factor subunit B (CBF-B); NF-Y protein subunit A (NF-YA); Hap2; CAAT-box DNA-binding protein subunit A	1.45	AAA40889.1
transcription factor LSF	1.37	B53771
homeobox 2.1 protein (HOX2A); HOXB5; HU1; HHO.C10	1.35	M92299.1
endothelial transcription factor GATA2	1.34	M68891.1
transcription factor Sp1 (TSFP1)	1.30	XM_028606.2
transcription factor ZFM1	0.62	G02919
zinc finger protein 161 (ZNF161); putative transcription activator DB1	0.26	NP_009077.1
stem cell protein (SCL); T-cell leukemia/lymphoma-5 protein (TCL5); T-cell acute lymphocytic leukemia-1 protein (TAL1)	-	AAA36598.1
neural retina-specific leucine zipper protein (NRL)	-	NP_006168
MSX-1 homeobox protein; HOX7	-	P28360
basic transcription factor 62-kDa subunit (BTF2)	-	AAA58399.1
paired box homeotic protein (PAX8) isoforms 8A/8B + isoforms 8C/8D	-	BAB59039.1
brain-specific homeobox/POU domain protein 3A (brn-3A); RDC-1; octamer binding transcription factor 1 (OTF1)	-	AAA65605.1
transcription factor E2-alpha (E2A); immunoglobulin enhancer binding factor E12; transcription factor-3 (TCF3)	-	AAA61146.1
transcriptional enhancer factor (TEF1); protein GT-IIc; transcription factor 13 (TCF13)	-	P28347
thioredoxin peroxidase 2 (TDPX2); thioredoxin-dependent peroxide reductase 2; proliferation-associated gene (PAG); natural killer cell enhancing factor A (NKEFA)	-	Q06830

Table 3 e: Receptors, Cell Surface Antigens, Cell Adhesion

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
interleukin-2 receptor gamma subunit (IL-2R gamma; IL2RG); cytokine receptor common gamma chain precursor; p64	*	AAA59145.1
interferon gamma receptor (IFNGR)	*	NM_000416.1
interleukin-1 receptor type I precursor (IL-1R1); IL-1R-alpha; p80; CDW121A antigen	*	M27492.1
neural-cadherin precursor (N-cadherin; NCAD); cadherin 2 (CDH2)	*	L34059
neural cell adhesion molecule L1 precursor (N-CAM L1); MIC5	*	M77640
integrin alpha 3 (ITGA3); galactoprotein B3 (GAPB3); very late antigen 3 alpha subunit (VLA3 alpha); CD49C antigen	*	M59911.1
leukocyte adhesion glycoprotein p150, 95 alpha subunit precursor; leukocyte adhesion receptor p150, 95; CD11C antigen; leu-M5; integrin alpha X (ITGAX)	*	M81695.1
integrin beta 4 (ITGB4); CD104 antigen	*	X51841.1
CD44 antigen precursor (CD44); phagocytic glycoprotein I (PGP1); HUTCH I; extracellular matrix receptor III (ECMR III); gp90 lymphocyte homing/adhesion receptor (LHR); hermes antigen; hyaluronate receptor; heparan sulfate proteoglycan; epican	1.51	XP_030326.1
glutamate receptor subunit epsilon 3 precursor (GRIN2C); N-methyl D-aspartate receptor subtype 2C (NMDAR2C; NR2C)	1.44	NP_000826.1
CD27L antigen receptor precursor; tumor necrosis factor receptor superfamily member 7 (TNFRSF7); T14	0.7	P26842
integrin alpha L (ITGAL); leukocyte adhesion glycoprotein alpha subunit precursor; leukocyte function-associated molecule 1 alpha chain (LFA1); CD11A antigen	0.45	P20701
interleukin 2 receptor alpha subunit precursor (IL-2 receptor alpha subunit; IL2RA); TAC antigen; CD25 antigen	0.41	P01589
CDW40 antigen; CD40L receptor precursor; nerve growth factor receptor-related B-lymphocyte activation molecule	0.35	CAA43045.1
granulocyte colony stimulating factor receptor precursor (GCSF-R); CD114 antigen	-	Q99062
low-affinity nerve growth factor receptor (NGF receptor; NGFR); GP80-LNGFR	-	AAB59544.1
neuromedin B receptor (NMBR); neuromedin-B-preferring bombesin receptor	-	NP_002502.1
granulocyte-macrophage colony-stimulating factor receptor alpha (GM-CSFR-alpha); CSW116 antigen	-	Q00941
platelet membrane glycoprotein IIIA precursor (GP3A); integrin beta 3 (ITGB3); CD61 antigen	-	P05106
integrin alpha 7B precursor (IGA7B)	-	CAA52348.1

Table 3 f: Growth Factors, Cytokines, Hormones

<u>Gene</u>	<u>Fold change</u>	<u>GenBank ID</u>
Interleukin-10 precursor (IL-10); cytokine synthesis inhibitory factor(CSIF)	**	M57627
Granulocyte-macrophage colony stimulatng factor (GM-CSF); CSF2	*	AAA52578.1
FMLP-related receptor I (FMLPRII); RMLP-related receptor I (RMLPRI)	*	AAA58482.1
Glia maturation factor beta (GMF-beta)	*	P17774
Hepatoma-derived growth factor (HDGF)	*	P51858
Macrophage inflammatory protein 1 alpha precursor (MIP1-alpha); tonsillar lymphocyte LD78 alpha protein; G0S19-1 protein; PAT 464.2; SIS-beta; small inducible cytokine A3 (SCYA3)	*	P10147
Monocyte chemotactic protein 1 precursor (MCP1); monocyte chemotactic and activating factor (MCAF); monocyte secretory protein JE; monocyte chemoattractant protein 1; HC11; small inducible cytokine A2 (SYCA2).	*	P13500
Oncostatin M (OSM)	*	NP_065391.1
Renin-binding protein (RENBP; RNPB)		XP_013053.3
Calgranulin B (CAGB); migration inhibitory factor-related protein 14 (MRP14); leukocyte L1 complex heavy chain; S100 calcium binding protein A9 (S100A9)	1.49	B31848
Placenta growth factors 1+2 (PLGF1 + PLGF2)	1.42	CAA38698.1
Vascular endothelial growth factor precursor (VEGF); vascular permeability factor (VPF)	1.42	AAA35789.1
Hepatocyte growth factor activator (HGF activator)	1.40	BAA74450.1
Follistatin-related protein precursor	1.34	AAA66062.1
Hepatocyte growth factor-like protein; macrophage stimulating protein (MSP)	1.29	AAA59872.1
Interferon gamma precursor (IFN-gamma, IFNG); immune interferon	1.29	P01579
WSL protein + TRAMP + Apo-3 + death domain receptor 3 (DDR3)	0.69	AAB41432.1
Neurotrophin-4 (NT4)	0.68	AAA60154.1
Interleukin-13 precursor (IL-13); NC30	0.39	P35225
Small inducible cytokine A5 (SYCA5); regulated on activation normal T-cell-expressed & secreted protein precursor (RANTES); SIS delta	0.38	XP_035842.1
Estrogen sulfotransferase (STE; EST1)	-	CAA72079.1
Keratinocyte growth factor (KGF); fibroblast growth factor 7 (FGF7)	-	AAA63210.1
Endothelial-monocyte activating polypeptide II (EMAP II)	-	AAA62202.1
Leukemia inhibitory factor precursor (LIF); differentiation-stimulating factor (D factor); melanoma derived LPL inhibitor (MLPLI); HILDA	-	B36282
Acidic fibroblast growth factor (AFGF) + heparin-binding growth factor 1 precursor (HBGF-1) + beta-endothelial cell growth factor (ECGF-beta)	-	AAA51672.1
Insulin-like growth factor-binding protein 3 precursor (IGF-binding protein 3; IGFBP3; IBP3)	-	P17936

Symbols (Fold Changes)

- 5 ** : Expressed in PP but not NPP, or in Co-culture but not Caco2.
 * : Expressed in Co-culture but no Caco2 (only repeated once)
 - : Expressed in Caco2 but not in co-culture.

Immunity

10 The events of the cell cycle occur under normal circumstances in a fixed sequence. Traditionally, the cycle is divided into two stages: cell division and the interphase. Cell division or mitosis is followed by cytokinesis and together they constitute the 'M phase' of the cell cycle. The interphase is divided up into the S, G₁ and G₂ phases. Briefly, during the S phase, DNA is replicated in preparation for mitosis, while the intermediate G phases are transitional periods involved in protein synthesis and cell growth. Activation of regulatory genes that control and maintain a cell's proliferative state by intracellular signals (discussed below) stimulates proliferation of the cell and initiates cell growth. A number of genes involved in these processes were differentially expressed in the co-culture model (as estimated by relative mRNA abundance) and discussed below.

20 The epithelial cells of the gut play an important part of the innate and specific immunity. IEC's are considered to be in a continuous controlled state of "physiological" inflammation and active processes continually take place to ensure that the tone of immunosuppression is maintained (Mayer, 2000). These unique regulators appear to control the mucosal immune system's condition. These distinct factors govern the immune response, whether it's immune suppression/tolerance, inflammation or a systemic immune response. A clearer understanding of the immunoregulatory features involved in mucosal immunity is clearly desirable and may lead to new approaches in disease and drug therapy. Genes detected in the co-culture model that may be related to or are involved in immune function in GALT are discussed below.

30 The gamma subunit of IL-2 receptor plays a pivotal role in formation of the full-fledged IL-2 receptor (Di Santo *et al.*, 1995). In an interesting study where infant rats were studied from pre- to post weaned life Masjedi *et al.* (1999) assessed alterations in expression and phenotype of cells in the gut-associated lymphoid tissue. At an age when the immune system is believed to be immature and functionally naive they discovered interleukin-2 receptor (IL-2R) expression peaked approximately four-fold at midweaning in Peyer's patches, compared with adult animals (day 70) suggesting that IL-2R expression is an adaptation to the host's environment. In a similar way, the presence of IL-2R specific for cells in the co-culture could be a direct result of the environment. The common gamma c

chain of the interleukin 2 receptor, gamma is also a component of the receptors for IL-4, IL-7, and IL-9 and plays a critical role in lymphoid development through its participation in the receptors for IL-2, IL-4, IL-7, IL-9, and IL-15 (Di Santo *et al.*, 1995)

Interferon- γ (IFN- γ) exhibits various properties including antigrowth activity in neoplastic and normal cells, and regulatory roles in immune responses (Tsuji *et al.*, 1998). Kjerrulf *et al.* (1997) found that in IFN- γ receptor knockout mice (IFN- γ R^{-/-}) reduced mucosal antibody responses and decreased Th1 and Th2 activity after oral immunization. The presence of IFN- γ receptor in the M cell co-culture model could possibly augment a cross-regulation between the two Th subsets in the gut mucosa. It is noteworthy that the ligand, IFN- γ , mRNA was increased in the co-culture that was supported further by the significant secretion of IFN- γ from co-culture monolayers.

The C-C chemokines macrophage inflammatory protein 1 (MIP1 α) and monocyte chemotactic protein (MCP1) are synthesized and expressed by epithelial cells (Vainer *et al.*, 1998; Kolios *et al.*, 1999). The purpose of these chemokines expression in the co-culture model could be to function not only in leukocyte migration, but also as adhesins in the interaction between leukocytes and colonic epithelium. However, another C-C chemokine, RANTES, mRNA was observed to be reduced in the co-culture. The reasons for this are unclear. Perhaps, the chemoattractant activities of other chemokines such as IL-8, MIP1 α and MCP1 are sufficient for the M cell and in the absence of T cells the need for RANTES is not required.

From a gene delivery perspective, a higher capacity for translation and protein synthesis in PP tissue indicates that PP tissue is a preferred tissue to which to deliver genes coding for DNA vaccines or antigens. Thus the proposed higher translational capacity of PP tissue has implications for gene delivery especially DNA vaccine delivery and correspondingly antigen expression and local presentation to the mucosal immune system in the gastrointestinal tract. The TF coding genes may be important in priming M cells or precursor cells to M cells to adopt M cell phenotype and/or to facilitate priming of M cells to give a better immune cell outcome.

M cell receptors identified in Table 3(e) above are of particular interest in that they can be used for vaccine and delivery.

In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of an IL-2 receptor, a gamma c chain of an IL-2 receptor, interferon - γ , and a C-C chemokine.

Proliferation and Growth

5 Cyclin D1 is a protein involved in regulation of the cell cycle. Over-expression of the protein is associated with abnormal growth or neoplasia. This protein is positively induced by the p42/p44 MAP kinases (Lavoie *et al.*, 1996). It would be interesting if the neoplasia seen in M cells resulted from activation of this protein considering the coincidental induction of the p44 MAP kinase (ERK1) below. The reduction in cyclin-dependent kinase 4 inhibitor 2D (CDKN2D) mRNA that normally inhibits cell cycle progression (Guan *et al.*, 1996) would insinuate a similar function in the proliferation of these 'M cells.'

10 In contrast, the induction of cell cycle inhibitors such as cyclin-dependent kinase inhibitor (CDKI) and cyclin-dependent kinase 4 inhibitor (CDK4J) would appear to be working to counterbalance proliferative stimuli present in the M cell.

15 PLC-L (phospholipase C-deleted in lung carcinoma) is a putative tumor suppressor gene. It is believed that irregular (in fact deletion) expression of the PLC-L gene contributes to the growth of human lung carcinoma (Kohno *et al.*, 1995). It is possible then that its upregulation in the M cell model is acting as a negative regulator of growth in the cells, counterbalancing the many proliferative signals present.

Growth factor receptor-bound protein 2, GRB2, involved in growth factor control of ras signalling (Lowenstein *et al.*, 1992).

20 The intracellular signaling pathways responsible for cell cycle arrest and establishment of differentiated cells along the gut axis remain largely unknown particularly in the case for the development of M cells and the FAE. ERK3/MAPK6 is expressed solely in the co-culture. Extracellular signal-regulated kinases-1 (ERK1) also known as the p44 mitogen-activated protein (MAP) kinase (p44mapk) is also induced specifically in the co-culture model. ERK1 and ERK3 are proline-directed serine/threonine kinases that are
25 activated in response to a variety of extracellular signals, including growth factors, hormones and, neurotransmitters. These MAP kinases are key molecules involved in intracellular signal transduction, and are key regulators of cell proliferation in mammalian cells (Davis, 1995). Results indicate that elevated p42/p44 MAPK activities stimulate cell proliferation of
30 intestinal cells, whereas low sustained levels of MAPK activities have correlated with cell cycle arrest and an increased expression of sucrase isomaltase (Aliaga *et al.*, 1999). It is tempting to speculate that the presence of ERK3 together with the other MAP kinases apart from their proliferative effects are in part responsible for a reduction in sucrase isomaltase, a characteristic effect in M cells.

Lying upstream in the ERK signal cascade the tyrosine/threonine protein kinase, MAPK kinase (MAPKK1) is implicated in the regulation of cell growth and differentiation through the activation of ERK. In addition it is interesting to note that MAPKK3 was deleted in the co-culture cells. MAPKK3 phosphorylates and activates p38 MAP kinase alpha and gamma isoforms (Enslen *et al.*, 1998). The induction of the MAPKK1 gene along with serine kinase coincides with the induction of ERK1, highlighting the ERK cascade as an important signalling cascade in M cell maintenance. It is interesting to note that ERK activation is responsible for terminal differentiation of components of the crypt-villus. (Taupin and Podolsky, 1999)

However, glia maturation factor- β (GMF- β) is potentially offsetting the ERK cascade effects. It is known to inhibit MAP kinases particularly ERK1 and ERK2 and yet promotes the p38 MAPK (Zaheer and Lim, 1996 and 1998).

Findings suggest that positive and negative regulation of MAPK activity are associated with loss of normal growth control and may be involved in carcinogenesis of colon cancers. Jun kinases such as JNK2 (MAP kinase 9) mediate signal transduction of pro-inflammatory cytokines and cellular stress (Uciechowski *et al.*, 1996).

CD40 is a receptor on the surface of B-lymphocytes, the activation of which plays critical role in B cell proliferation and differentiation. CRAF1, (CD40 receptor-associated factor 1) encodes a protein that interacts directly with CD40 receptor (Cheng *et al.*, 1995). Its upregulation in the co-culture is perhaps a main determinant of lymphoepithelial crosstalk as discussed above.

The c-myc gene is commonly amplified and over-expressed in many human tumors (Ryan and Birme, 1996). A member of the myc family of helix-loop-helix transcription factors, c-myc is integral in controlling cell growth and promotes cell proliferation and transformation by activating growth-promoting genes (Thompson, 1998). Prothymosin- α (PT- α) is a nuclear protein and its expression is associated with alterations in the proliferative state of cells and has been reported to be regulated by the c-myc gene in vitro. (Smith, 1995; Mon *et al.*, 1993). The increased activity of c-myc in this model is likely to result in the increase in RT- α mRNA.

PKC- α protein levels regulate certain pathways that lead to the expression of differentiation-dependent genes. In a series of antisense transfection experiments where PKC- α expression in CaCo-2 cells was almost completely deleted, enhanced proliferation and a marked decrease in differentiation was observed, as well as a more aggressive transformed phenotype (Scaglione-Sewell *et al.*, 1998). In a similar fashion, the lack of PKC-

α mRNA detected in the co-culture 'M cells' may underlie some of the phenotype changes featured.

5 Glutathione S-transferase A1 (GSTA1) is a member of a multigene family of detoxification and metabolizing enzymes. Induction of GST enzyme activity has been demonstrated to act as a potent anti-proliferative and differentiating agent in Caco-2 cells (Stein *et al.*, 1996) suggesting a similar role in the 'M cell.'

 Transcription factor GATA-2 is thought to maintain and promote the proliferation of early haematopoietic progenitor cells.

10 The placenta growth factor (PLGF) is a member of the vascular endothelial growth factor (VEGF) family of growth factors. In addition to PLGF, VEGF mRNA was enhanced in the co-culture cells. These growth factors play a crucial role in angiogenesis during development and/or repair (Andre *et al.*, 2000). The augmented transcription of their mRNA is consequently not a surprising find. However, hypoxia and energy depletion are known to induce angiogenesis by increasing VEGF, expression and so the possibility that the co-
15 culture conditions are responsible for these genes induction cannot be ruled out rather than a deliberate mechanism of neogenesis in M cell formation. VEGF receptor 1 (VEGFR1); the receptor for VEGF and PLGF, mRNA is down-regulated and is possibly a consequence of desensitization of the receptor by VEGF and PLGF binding, initiating a reduction in the receptor's RNA.

20 Coinciding with the above actions, the absence of growth factors such as insulin-like growth factor-binding protein 3 (IGFBP3) and keratinocyte growth factor (KGF) may be modulating enterocytic cell proliferation and differentiation.

 Caco-2 cells have been shown to express the type I IL-1R. (Varilek *et al.*, 1994) IL-1R α binds IL-1 and mediates cell signalling particularly signalling involved in cell proliferation
25 (French *et al.*, 1996). The expression of IL-1R can be enhanced by IFN- γ (Varilek *et al.*, 1994). Therefore, the expression of IL-1R type 1 mRNA in the co-culture is interesting when considering the significant expression of IFN-1 expressed in supernatants of the co-culture model.

30 In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, PKC- α , GSTA1, GATA-2, and PLGF.

Differentiation

Development of cells or differentiation is dictated by the expression of a cell's genes specific to that cell. This is a particularly important aspect with regards to M cells.

5 The cortical cytoskeleton not only provides structural support to the plasma membrane but also contributes to important dynamic processes such as endocytosis, exocytosis, and transmembrane signalling pathways. Ezrin, or villin 2, is an F-actin associated molecule and is concentrated in surface projections such as microvilli and membrane ruffles where they link the microfilaments to the membrane and has been
10 reported to be in abundance during development and differentiation of the intestinal epithelium. It was reported that hepatocyte growth factor (HGF/SF) could stimulate the tyrosine phosphorylation of ezrin in a human colon epithelial cell line, which induced the ezrin associated membrane ruffling. It is interesting to note that both hepatocyte growth factor activator (HGF activator) and hepatocyte growth factor-like protein were both upregulated in
15 the co-culture model and taken with the augmented ezrin mRNA the induction of these genes would appear to underlie the mechanism involved in the morphogenesis observed in M-cells.

 These data demonstrate that the expression of the ezrin gene is being regulated at the level of mRNA due to effects incurred by the B-cells. It is particularly relevant
20 considering the observations of villin diffusely displayed in M-cells.

 One method of actin cytoskeletal reorganization is controlled by the LIMK-1 serine/threonine kinase, which acts by phosphorylating cofilin and subsequently Rac (as previously reported). However, LIMK-1 was deleted in the co-culture model and would appear to rule out the Rac-mediated mechanism of actin reorganization in the M cell model.

25 The cadherin family of cell adhesion molecules play important role in cell-cell adhesion during tissue differentiation. They have been reported to be linked to the actin cytoskeleton by catenins located in the cytoplasmic compartment of the cell. The specific expression of NCAD in the co-culture suggests a distinct gene involved in the cytoskeletal structure.

30 Previous reports have shown that neogenin is closely related to the human tumor suppressor molecule DCC (deleted in colorectal cancer) and together they constitute a subgroup of Ig superfamily proteins that have shown to be essential for terminal differentiation of specific cell types in the adult including the human colon. These parallels suggest that neogenin, like DCC, is functionally involved in the transition from cell

proliferation to terminal differentiation of specific cell types. Its absence in the co-culture model might represent a period of continued proliferation for the cells and allow a longer period of proliferation.

5 The helix-loop-helix (HLH) family of transcription factors has been shown by others to play a central role in the regulation of cell growth, differentiation and tumorigenesis. Of particular interest, when HLH 1R21 was over-expressed in mouse NIH3T3 cells, it induced a morphologically transformed phenotype.

10 Other genes associated with differentiation including Myeloid cell nuclear differentiation antigen (MNDA) and the LHX1 gene. The LHX1 gene is a member of the LIM/homeobox (Lhx) gene family. It has been shown that it codes for a transcriptional regulatory protein involved in the control of differentiation and development.

15 In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of HGF activator, ezrin, NCAD, MNDA, and LHX1.

Adhesion

20 It is clearly evident that modification of the M cell apical surface is a determining factor in M cell apical membrane adherence, and thus, uptake and transport of macromolecules/microorganisms and targeting epitopes on the surface of M cells has been used to promote further adherence and uptake of particles in vaccinology. The specificity of these markers is not only useful for vaccine strategies but also represents targets for
25 understanding adhesion and uptake of bacteria and viruses. Adhesion is not privy to the apical surface. Adhesion molecules on the basolateral surface of M cells, such as cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4 are understood to be involved in leukocyte migration and in the development/organization of lymphoid nodules in Peyer's patches. Genes
30 expressed/induced in the co-culture can provide an insight into the mechanisms involved and are discussed below.

The tyrosine kinase receptor TIE 1 is normally located in vascular endothelial and haematopoietic cells and is largely involved in the proliferation and differentiation of miniature haematopoietic cells and would be an appropriate gene specific for M cells. In the brain, TIE

mRNA and protein is significantly elevated in lesions composed of abnormal vasculature called arteriovenous malformations (AVMs) and the surrounding vasculature. Like AVMs, the significant upregulation of TIE in M cells may indicate some ongoing neogenesis, and depending on the receptor's polarity could be of potential use in vaccine targeting.

5 The neuronal cell adhesion molecule L1 (NCAML1) is a transmembrane glycoprotein belonging to the immunoglobulin superfamily and is generally associated with development of the nervous system. As a potent promoter of neurite growth, it is allied with plastic changes. In nerve growth it interacts with the actin cytoskeleton via an ankyrin linkage and promotes specific distribution of F-actin. Such flexibility is ideal in the M cell scenario.

10 The integrin family consists of a series of related alpha beta heterodimers involved in a variety of cell-matrix and cell-cell adhesion functions. The $\alpha_3\beta_1$ integrin is a multiligand extracellular matrix receptor found on many cell types and can function as a receptor for fibronectin, laminin, and collagen. Phagocytosis of molecules by breast cells has also been reported to involve this adhesion molecule, thus, it would appear a suitable candidate as an

15 adhesion target on M cells.

 The leukocyte adhesion glycoprotein p150 (CD11C antigen), also a member of the integrin family, is involved in leukocyte sequestration via interaction of CD11/CD18 similar to that of ICAM-1.

20 In stratified epithelia β_4 integrin (CD104 antigen) has been shown to be important for proper differential expression and crucial for stable adhesion to the basement membrane through its ability to attach externally to laminin and internally to the keratin cytoskeleton. Interestingly, during human intestinal organogenesis receptors have been shown to occur. This integrin would appear to play an important role in epithelial cell-matrix interactions during development but particularly in M cell development.

25 CD44 is a major surface adhesion molecule involved in cell-cell and cell-matrix interactions and lymphocyte homing and activation. The observed enhanced expression suggests that this molecule is an important feature in the activities of M cells. A non-receptor tyrosine kinase, C-src protooncogene (SRC1) has been shown to cause overexpression of CD44 in the intestine. As well as its effects on proliferation, the enhanced

30 activity of SRC1 seen in the M cell model would appear to have major effects on cell adhesion properties of the M cell. Hepatocyte growth factor activator (HGF activator) is a serine protease produced and normally secreted by the liver. It has been documented as stimulating reparative processes in intestinal epithelial cells and could be why its activity is enhanced in this model. However, stimulation of CD44 in colonic epithelial cells has been

reported to augment c-met, the HGF receptor. This in turn stimulates the "inside-out signalling causing an amplified expression of integrins that leads to an increase in vascular adhesion to the epithelium.

5 It has been reported that the glutamate receptor (NMDA) is generally associated with learning and memory, highly plastic processes in the brain. The high density of NMDA receptors reflects similar plastic changes seen in the co-culture model but would also act as a target epitope for drug delivery.

10 TKT is a tyrosine-kinase receptor related to TRK and is a member of cell adhesion kinase receptor family. Ephrin (type A) is a tyrosine kinase receptor that has been reported to be involved in neogenesis and tumor formation. Sp1 is a nuclear protein constitutively expressed and mediates basal promoter activity and is the main Vitamin-D receptor promoter in intestine. These are all potential target sites relevant to M cells.

15 Many of the receptors/cell surface antigen 'deleted' (not detectable) in the co-cultures could be putative negative markers of M cells. A good example is the laminin receptor $\alpha_7\beta_1$ integrin. Expression of the $\alpha_7\beta_1$ integrin correlates with human intestinal cell differentiation and could be used in a similar fashion that was applied with sucrase isomaltase and alkaline phosphatase.

20 In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, $\alpha_3\beta_1$ integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1.

25 Transport

30 The RAB proteins are reported to be regulators of polarized membrane traffic in epithelial cells. The RAB3B is localized to the apical pole very near the tight junctions between adjacent epithelial cells where it is reported to be a possible regulator of apical and/or junctional protein traffic in epithelial tissues. RAB3B is highly homologous to a brain-specific RAB3 isoform (RAB3A) that targets the presynaptic nerve terminal, where it is reported to regulate exocytosis.

In polarized cells, the small GTPase Rab5a is localized to the plasma membrane, clathrin-coated vesicles, and early endosomes and is a regulator of transport between the

plasma membrane and early endosomes. The decreased expression of RAB5a seen in the co-culture may deregulate the rate of endocytosis and/or vesicle fusion and could possibly release 'the brake' on vesicle trafficking.

5 RAB6 is another ras related protein also a regulator of intracellular transport in mammalian cells. It controls intra-Golgi transport, either acting as an inhibitor in anterograde transport or as a positive regulator of retrograde transport. Like RAB5a, the pronounced decrease seen in mRNA transcription could be a means of subverting transport regulation in epithelial cells and so optimize the process as observed in M cells.

10 Protein kinase C (PKC) and the actin cytoskeleton are critical effectors of membrane trafficking in mammalian cells. The F-actin cross-linking protein myristoylated alanine-rich C kinase substrate (MARCKS), a substrate for PKC, has been reported to be a component of the mechanism of endocytosis.

15 TIR or p71 plays a key role in the control of cell proliferation through the binding of transferrin, the major iron-carrier protein. Located on both apical and basolateral surfaces, the transferring receptor has the ability to internalize and recycle to the surface. Indeed experiments by Hughson and Hopkins (1990) demonstrate pathways from the apical and basolateral surfaces meet in an endosomal compartment. Furthermore, Shah and Shen (1994) discovered that the fungal metabolite brefeldin A (BFA) could relocate receptor distribution and enhance TfR mediated transcytosis. The increased expression of this
20 mRNA in the M cell model suggests a potential delivery mechanism of protein drugs across the intestinal epithelium present in M cells that could be exploited.

In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of a RAB protein, PKC, and TfR.

25

Signal transduction

30 In order for a cell to respond to extracellular signals, which cause it to alter gene expression or cellular function, it must involve the activation of a signal transduction cascade. There are many different types of signalling cascades, which can be unique to a specific type of stimulus. There are two main mechanisms by which these cascades transmit their signal, either through the regulation of enzymes, which produce second messenger molecules or through the regulation of protein phosphorylation. The activation of these cascades is usually mediated through specific cell surface or intracellular receptor proteins. The receptor

protein recognizes the incoming extracellular signal and responds accordingly, initiating a specific series of intracellular signal that direct the cell's behavior. A number of genes involved in intracellular signalling were upregulated or induced in the M cell model and are discussed below.

5 A member of the Janus family of tyrosine kinases, which are non-receptor protein kinases, Jak3 is involved in intracellular signalling mediated by cytokines and growth factors such as IL-2, IL-4, and IL-7. Jak3 has been reported to play a crucial role in Peyer's patch organogenesis. Mutant mice deficient in Jak3 presented defects in lymphocyte production and the absence of Peyer's patch structures. Its induced expression suggests a greater
10 level of activity and possibly a major requirement underlying the M cell phenotype 'switch'.

 The nuclear zinc-finger transcription factor, early growth response factor-1 (EGR-1) is an immediate-early gene product expressed in response to diverse stimuli and is involved in growth, development, and differentiation. EGR-1 has been reported to function in growth regulation and suppression of cell transformation by transactivation of the TGF β gene.
15 TGF β is capable of stimulating the synthesis of extracellular matrix proteins that can potentially stabilize epithelial cell contact with the substratum. In addition EGR-1 also plays a role in the immune response, regulating targets such as IL-2, CD44, ICAM-1, and TNF. Taken together the considerable induction of EGR-1 mRNA emphasizes the importance of this protein's involvement in M cell behavior.

20 CaM kinase IV (CAMK IV) is involved in Ca²⁺-dependent mechanism for regulating MAP kinase pathways. Many kinases activity has been observed to be enhanced in this model and so it is logical that CAMK IV expression is induced as a requirement to function.

 The tyrosine kinase Tnk1 has been reported to be involved in signalling pathways involving development in adult tissues and in cells of the lymphohaematopoietic system.

25 Epithelial discoidin domain receptor 1 (EDDR1) mRNA was reduced in the co-culture. EDDR1 is a collagen receptor involved in controlling cellular responses to the extracellular matrix (ECM). The decrease in this gene would implicate it in the reorganization of the M cell in relation to the ECM.

 cAMP-dependent protein kinase type I beta regulatory subunit (PRKAR1B) stimulates
30 growth by modulating the signalling of camp via its regulation of cAMP-dependent protein kinase (PKA). PRKAR1B's reduction in the co-culture model may represent an inhibitory role in the cell's growth counterbalancing the proliferative signals.

 In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the

protein may be selected from the group consisting of Jak 3, EGR-1, TNK1 and CAMK IV.

Protection and repair

5 Chaperones such as HSP40 and HSP 70 participate in many biological processes in which protein folding is involved. These include protein translocation, protein translation, protein assembly and disassembly, and protein degradation. It is understandable that such genes would be induced considering the evolving processes of a phenotype 'switch.' However heat shock protein production has been reported to be induced as a result of harsh
10 changes in their environmental conditions such as stress, ischaemia or hypoxia resulting in protein damage. Therefore it cannot be ruled out that the induction of these genes is in fact a protective measure as a consequence of the adverse conditions in the co-culture.

 HSP 60 has been observed in highly replicating cells e.g. short-living epithelial cells of the intestine. Involved in the import and refolding of nuclear-encoded proteins destined for
15 the mitochondrial matrix.

 The 27-kDa heat shock protein (HSP27) is expressed in a variety of tissues, including gut epithelia and in the absence of stress has been reported to regulate actin filament dynamics. Hsp27 induction in the M cell model like the other heat shock proteins (HSPs) may be active in development of resistance to stressful conditions. Activation of HSP27 can
20 contribute to agonist-induced phosphorylation-modulated reorganization of the actin cytoskeleton and, in the case of stress activation, provides an actin-based adaptive response of cells to the new environmental conditions, and is ideal candidate for the plasticity seen in M cells.

 Expression of receptors for fMLP on human phagocytes is well established, but there
25 is conflicting evidence regarding the potential expression of fMLP receptors on other cells within the mucosa, particularly the epithelial cells. The reported observation of the receptor for the chemotactic peptide fMLP supports the notion of the intestinal epithelial cell as an early "sensor" of infection and inflammation. It has been reported that, fMLP, present in abundance in the lumen of the gut and that activation of fMLP receptors induces cytotoxic
30 effects such as lysosomal release and superoxide generation. Thus, it would appear that their presence would be a defensive role in the event of infection of microorganisms.

 Glutaredoxin (thioltransferase) is a small, heat-stable protein catalyzing glutathione-dependent disulfide oxidation reactions in a coupled system with NADPH, GSH and glutathione reductase. It is important in regulating cell metabolism through the inactivation of

oxidated transcription factors thought to be important in cellular responses to oxidant stress. This modulation of transcription factors' binding activity has been demonstrated for a number of transcription factors, including NF-kB/Rel proteins, Fos and Jun proteins and nuclear factor I (NFI) family of transcription factors. The induction of such a gene would appear to provide a protective role and is particularly influential on a number of key transcription factors.

CREB has been implicated as having prominent role in protection. Over-expression of the gene was reported to reverse hypoxia elicited TNF induction. This infers that the increase in the cAMP responsive element binding protein (CREB) mRNA is possibly a protective response to conditions.

Inactive in cells under normal conditions, gadd153 expression is markedly induced in response to a variety of cellular stresses, including nutrient deprivation, DNA damage, and oxidative stress (e.g. free radicals) which normally leads to growth arrest. The arrest in growth is thought to allow critical repair processes to be carried out before any further cell cycling. It would appear that the gadd153 expression in the co-culture is for reparative purposes.

The excision repair proteins XPG and XPD have been reported to be involved in nucleotide repair. In addition, mRNA for ubiquitin-conjugating enzyme (likely to be involved in post-replication repair and induced mutagenesis, RAD23, and ataxia telangiectasia are also expressed in the co-culture. Their expression, coinciding with gadd153 suggests there is a high degree of impairment to genes in the M cell model.

Interleukin-13 (IL-13) is a potent anti-inflammatory cytokine and has been reported to have the same protective properties in inflammation as IL-4 through its ability to modulate and suppress pro-inflammatory cytokines. It is puzzling that in an environment with a high level of pro-inflammatory cytokines produced that IL-13 mRNA is in fact reduced. One possible explanation might be its anti-adhesion effect. It has been reported that IL-13 (secreted from lymphocytes) down regulated cell adhesion molecules in colonic epithelium and so the role of IL-13 in the co-cultured cells is modulating cell adhesion properties and not inflammation.

In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group consisting of HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd 153, XPG, XPD, ubiquitin, conjugating enzyme, RAD 23, and ataxia telangiectasia.

Apoptosis and programmed cell death

In programmed cell death, apoptosis is programmed in the sense that a genetically directed 'clock' selects a given time for the death of certain cells. It has been reported that it provides an important mechanism for the maintenance and renewal of cells in the gut and in development. However, for the epithelium to maintain its barrier functions, the level of apoptosis needs to be regulated, and this is 'checked' by several signal transduction systems. Toxic insult or lack of factors that maintain cell survival can also lead to apoptotic death of the cell.

It has been reported that over-expression of c-fos and c-jun (constituents of the AP-1 transcription factor) in the intestine correlates with programmed cell death and subsequent cellular regeneration. Other studies have demonstrated increases in both proximal jejunum and colon jun mRNA level coincide with a period of major changes in intestinal cell proliferation). The c-jun protein product involved in activation of AP-1, transcription is enhanced when it is phosphorylated by stress-activated protein kinases of which there are many in the M cell model.

As intestinal epithelial cells reach the villus apex they undergo apoptosis and, are shed and, in normal circumstances, caspases, a family of cysteine proteases, play a central role in initiating, amplifying, and executing apoptosis. The pattern of caspase activation in this process is not understood. It is interesting to note that the apoptosis regulator, bcl-x, and caspase 9 are induced in the co-culture. The bcl-x gene plays an important role in the regulation of programmed cell death (PCD), depending on its splice variant the bcl-x protein can accelerate apoptosis or delay/prevent programmed cell death (as previously reported). Bcl-x controls apoptosis mechanisms at points upstream of caspase activation. Perhaps, it is responsible for the marked induction of caspase-9. Caspase-9 is a caspase initiator. Once activated, it can proteolytically activate other caspases (including 3, 6 and 7), which in turn activate caspase-2 and 6 (as previously reported). Inhibitor of apoptosis protein 2 (IAP2) binds to and inhibits caspase-3. Its expression is a mechanism of regulating cell death depending on the particular cellular or environmental signals. Therefore, its absence in the co-culture cells and the increased activity of caspase-9 allows caspase-3 unchecked pro-apoptotic activity.

The death domain receptor 3 (DDR3) member of the TNFR family can induce apoptosis as previously reported. Its mRNA expression is also reduced in the co-culture model.

In view of the foregoing, in the method of the invention for increasing the level of a protein in a PP cell, which comprises delivering a nucleic acid coding for a protein, the protein may be selected from the group selected from: bcl-x and capase-9 and more generally in view of the foregoing may be selected from the group consisting of

5 cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, CD40, C-MYC, PKC- α , GSTA1, GATA-2, PLGF, ezrin, HGF activator, hepatocyte growth factor-like protein, NCAD, MNDA, LHX1, TIE-1, NCAML1, CD104, CD44, SRC1, NMDA, TKT, ephrin (type A), Sp1, RAB proteins, PKC, TIR, Jak3, EGR-1, TNK1, CAMK IV, HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin,
10 CREB, gadd153, XPG, XPD, ubiquitin- conjugating enzyme, RAD23, cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, α 3 β 1 integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1, a RAB protein, PKC, and TfR, bcl-x and capase-9

Example 6

15

Targeted Gene delivery

Delivery of genes, gene fragments, oligonucleotides or other nucleotide fragments or analogues of the present invention to a living organism can be accomplished by methods currently available in the prior art. For example, various recombinant viruses have been
20 used for the oral delivery of genes, such as adenovirus, retrovirus, adeno-associated virus, vaccinia virus, lenti-virus and plant-derived viruses, wherein the viral genome is replaced with an expression vector for the gene of interest. . See, David T. Page and Sally Cudmore (2001). Innovations in oral gene delivery: challenges and potentials. *Drug Discovery Today*, Vol. 6, No. 2, pp 92–101. Viral mimetic particles such as virosomes and various types of
25 polymers and liposomes, such as cationic and fusogenic, are also employed for gene delivery. See, U.S. Patent Nos. 4885172, 5047245, 5171578, 5059421, 5399331, 5204112, 1252263, 5376452, 5552155, 6120797, 6087325, 6143716. Examples of polymers are PLGA, PLA co-polymers, chitosan, and fumaric acid/sebacic acid co-polymers. For these systems, the polymer or liposome is formed from component parts in a solution of the gene
30 expression vector, thus encapsulating the genes when particles are formed. Cationic lipids such as DOTAP and polyethylenimine are commonly used whereby the gene expression vector is complexed with and protected by the lipids. (See, Ogris M. *et al.* (2001). DNA/polyethylenimine transfection particles: Influence of ligands, polymer size, and pegylation on internalization and gene expression. *AAPS PharmSci*, 3 (3), article 21).

Agents such as protamine are used to condense DNA, which due to the reduction in size of the DNA particles are more easily taken up by cells. Recombinant live bacteria (e.g. *Shigella* spp, *Salmonella* spp.) have also been exploited for gene delivery to the gut. Oral bioavailability enhancers, (e.g. sodium caprate, Elan's PROMDAS technology) could be used to increase uptake of a gene or encapsulated gene formulation.

In all cases the delivery systems can be targeted with various ligands on the surface of the particles in order to enhance binding to specific cells type and/or to enhance uptake. These ligands could be peptides, proteins, antibodies, peptidomimetics, and lipids that recognize or are being recognized by specific sites/receptors on the cell surface (Maruyama K. (2000). *In vivo* Targeting by Liposomes. *Biol. Pharm. Bull.*, 23(7), 791-799).

The targeting ligands may be peptide based, peptidomimetic based, antibody based, single chain antibody based, small organic molecule based. The targeting ligands may also be natural substrates for such receptors, transporters or other cell surface molecules found on the surface of M cells or other cell types found in Peyer's patch. The targeting ligands may be engineered so as to be genetically expressed on the surface of viruses, bacteriophages, virosomes, bacteria or other organisms, which can be utilized for vaccine delivery in the gut. Furthermore the targeting ligands can be presented either as direct conjugates to antigens, or on the surface of drug-loaded particulates such as liposomes, PLGA particles, other particulates and at the same time retain recognition by and interaction with the receptors, transporters or other cell surface molecules found on the surface of M cells and / or other cells of Peyer's patch tissue.

Examples of peptides that target the gastro-intestinal tract, in particular, membrane translocating peptides useful for vaccine delivery to M cells along with M cell specific targeting ligands are described in Table 4.

Further, targeting ligands can be genetically engineered into the surface coats of viruses, bacteriophages or bacteria, conjugated directly to antigens conjugated to the lipids in liposomes by covalent methods or streptavidin-biotin linkages, or coated onto the surface of polymers after particle formation (Torchilin V.P. *et al.* (2001) *Proc. Natl. Acad. Sci. USA*, Vol. 98, Issue 15, 8786-8791, July 17).

TAT peptide on the surface of the liposomes affords their efficient delivery even at low temperature and in the presence of metabolic inhibitors; Lestini *et al.* (2002). Surface modification of liposomes for selective cell targeting in cardiovascular drug delivery. *J. Controlled Release*, 78, 235-247; Dokka S. *et al.* (1997) Cellular delivery of oligonucleotides by synthetic import peptide carrier. *Pharm. Res.*, vol. 14, No. 12, 1759-1764; Wu Y *et al.*

(2000). Gene transfer facilitated by a cellular targeting molecule, retrovirus protein σ 1. *Gene Therapy*, 7, 61-69).

5 When the delivery of the gene to M cells in the gut is designed to prime or boost the immune system, the genes can be co-delivered/co-encapsulated with adjuvants (e.g. MF59, alum, saponin, QS21, MPL, bacterial toxins such as Lt, CT or mutants there-of, CPG motif nucleotides). Immune response could be boosted at a later stage by methods such as subcutaneous administration of an adjuvant.

10 In some cases it may be desired to shut off expression of certain genes, so as to enhance the adoption by enterocytes of an M cell phenotype. This can be achieved by the delivery, by methods outlined above, of antisense oligonucleotides, ribozyme, or RNA-interference molecules specific to the gene of interest.

Table 4

SEQ ID. NO:

SEQ ID. NO:	PEPTIDE SEQUENCES
SEQ. ID NO: SEQ. ID NO	GPHRRGRPNSRRSSKT GTSNGNGCCNYDGP
	<u>Peyers patch and/or M cell specific targeting ligands:</u>
SEQ. ID NO:	ATPPPWLLRTAP
SEQ. ID NO:	DGSIHKRNIMPL
SEQ. ID NO:	DYDSLWRSTLH
SEQ. ID NO:	GEPTTDMRWRNP
SEQ. ID NO:	GLWPWNPVTVLP
SEQ. ID NO:	HMLNDPTPPPYW
SEQ. ID NO:	KPAYTHEYRWLA
SEQ. ID NO:	LETTCASLCYPS
SEQ. ID NO:	LGTDWHSVSYTL
SEQ. ID NO:	LGTLNAGVPGFP
SEQ. ID NO:	LTHSKNPVFLST
SEQ. ID NO:	LVPTTHRHPVT
SEQ. ID NO:	LVSNARGFNNLS
SEQ. ID NO:	NTRIEPIRFYM
SEQ. ID NO:	NVYTFHSMSPMP
SEQ. ID NO:	QHTTLTSHPRQY
SEQ. ID NO:	SDFSDTMPHRPS
SEQ. ID NO:	SIDTIQILSLRS
SEQ. ID NO:	SISWASQPPYSL
SEQ. ID NO:	SMVKFPRPLDSR
SEQ. ID NO:	SPTLGASVAQTN
SEQ. ID NO:	TMSPNVYYTAFG
SEQ. ID NO:	TQIPSRPQTPSQ
SEQ. ID NO:	VCSNMYFSCRLS
SEQ. ID NO:	VPPHPMTYSCQY
SEQ. ID NO:	VPRLEATMVPDI
SEQ. ID NO:	VPTKPELPVNFT
SEQ. ID NO:	WSSDLPQPASTY
SEQ. ID NO:	YITPYAHLRGGN
SEQ. ID NO:	NVYTDNTLSPTP
SEQ. ID NO:	LETTAASLCYPS
SEQ. ID NO:	SPYCLSACTTEL
SEQ. ID NO:	LETTCASLCYPS
SEQ. ID NO:	VPPHPMTYSCQY
SEQ. ID NO:	VPPHPMTYSAQY
SEQ. ID NO:	VPPHPMTYSSQY
SEQ. ID NO:	YQCSYTMPPPV
SEQ. ID NO:	VCSNMYFSCRLS
SEQ. ID NO:	VSSNMYFSSRLS
SEQ. ID NO:	DYDSLWRSTLHGHESSH
SEQ. ID NO:	GNPTSTMW
SEQ. ID NO:	PWNSATVL
SEQ. ID NO:	NDPTAPPY

SEQ ID. NO:	PEPTIDE SEQUENCES
	<u>Membrane Translocating Peptides:</u> (underline denotes cyclization)
SEQ. ID NO:	KKAAVLLPVLLAAP FITC-LC
SEQ. ID NO:	KKKAAVLLPVLLAAP
SEQ. ID NO:	KKAAVLLPVLLAAPREDL
SEQ. ID NO:	<u>KKCAAVLLPVLLAAPC</u>
SEQ. ID NO:	<u>CAAVLLPVLLAAC</u>
SEQ. ID NO:	<u>KKCAAVLLPVLLAC</u>
SEQ. ID NO:	CAAVLLPVLLC
SEQ. ID NO:	CAAVLLPVLC
SEQ. ID NO:	CAVLLPVLLAAPC
SEQ. ID NO:	CVLLPVLLAAPC
SEQ. ID NO:	CLLPVLLAAPC
SEQ. ID NO:	CLPVLLAAPC
SEQ. ID NO:	AAVLLPVLLAAP .
SEQ. ID NO:	AAVLLPVLLAA
SEQ. ID NO:	KKAAVLLPVLLA
SEQ. ID NO:	AAVLLPVLL
SEQ. ID NO:	AAVLLPVL
SEQ. ID NO:	AVLLPVLLAAP
SEQ. ID NO:	VLLPVLLAAP
SEQ. ID NO:	LLPVLLAAP
SEQ. ID NO:	LPVLLAAP
SEQ. ID NO:	AAVLLPVLLAAKKKRKA
SEQ. ID NO:	KKKRKAAA VLLPVLLA

Example 7

5 Use of bacterial coatings to convert enterocytes to M cells

10 Might be nice to have some type of claim capturing the concept from this section Use
of bacterial coatings on PLGA particles, co-administered bacterial particles or pro-biotic
yogurts as adjuvants for oral vaccination with PLGA particles. The invention is based on
converting enterocytes to M cells by using specific bacteria in advance of, or along with the
oral vaccine particle of interest. In doing so the capability of absorbing particles through M
cells will be increased. This idea is not based on targeting but on the ability of live bacteria
or active bacterial components to stimulate cytokine production in Peyer's patches, thus,
enabling enterocyte-M cell conversion. As a result, an invention disclosed herein is a
method of promoting enterocyte-M cell conversion, said method comprising orally
15 administering an antigen, antigenic composition, or antigen-carrying particle to a person and
either simultaneously with, or prior to, said administration, also orally administering a

bacteria, or pro-biotic yogurts, or bacterial component to said person.

All references cited herein are incorporated herein by reference in their entireties.

5

Table 5
Miscellaneous GenBank Accession Numbers

Human Serum Albumin	NM_000477.3
Calreticulin	M84739

10

15

Dates for GenBank records

To the extent the date of a GenBank record, rather than its version number, is relevant for purposes of incorporation by reference, the date of the record is the filing date of this application with the following exceptions:

20

Table 2: Rat genes

3/27/02	D83697 through M10149
3/28/02	Q03238 through NM_017218.2

25

Tables 3: Human genes with a fold change of 0.5 or less
4/02/02

Table 2 Human genes with a fold change of 0.5 or less

4/02/02	U76376.1 through XM_087242.1
4/03/02	S90469 through M29366.1

30

The records specified for 3/27/02, 3/28/02, 4/02/02, and 4/03/02, do not include those

of GenBank IDs: Q07912, P21145, P46734, Q92851, Q13490, NP_006168, P28360, P28347, Q06830, P20701, P01589, P05106, P35225, P17936, S18408, P17074, P10661, P35426, P09456, P22791, Q06486, P21708, Q03238, P10644, P54868, P10398, P48730, P31749, P27361, P25063, Q14012, P13866

CLAIMS

1. A method of increasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell a nucleic acid coding for a protein, wherein absent
5 said increase, the levels of said protein or its mRNA is greater than in a non-Peyer's patch cell.
2. The method of Claim 1 wherein the protein is a transcription factor or a protein that activates a transcription factor.
3. The method of Claim 2 wherein the transcription factor or a protein that
10 activates a transcription factor is selected from the group consisting of Jun-B; c-jun related TF, Jun-D; c-jun related TF, STAT 3 - signal transducer and activator of transcription 3, NF-kappa β Tf p105 subunit, , S-myc proto-oncogene; myc related, Nm23-M2; nucleoside diphosphate kinase B; metastasis reducing protein, and C-est-I proto-oncogene; p54.
4. The method of Claim 1 wherein the protein is a receptor, or cell surface
15 antigen,
5. The method of Claim 4 wherein the protein is a receptor or a transporter.
6. The method of Claim 1 wherein the protein is selected from the group consisting of nucleoside diphosphate kinases and member of the 14-3-3 family.
7. The method of Claim 1 wherein the protein is coded for by a gene with an
20 expression Fold Change denoted by a **, *, or number greater than 2.00 in Tables 2 or 3.
8. The method of Claim 1 wherein the nucleic acid coding for at least 2 proteins is delivered, each of said proteins coded for by a gene with an expression Fold Change denoted by a **, *, or number greater than 2.00 in Tables 2 or 3.
9. The method of Claim 1 wherein the cell to which the nucleic acid is delivered
25 is a human cell.
10. The method of Claim 9 wherein the cell is in a Peyer's patch in a human and the nucleic acid is delivered by the oral route.
11. The method of Claim 9 wherein the cell is not within the body of a human.
12. The method of Claim 1 wherein the cell to which the nucleic acid is delivered
30 is a rat cell.
13. The method of Claim 1 wherein a nucleic acid coding for a tumor antigen or foreign peptide is also delivered to the Peyer's patch cell.
14. The method of Claim 13 wherein the cell to which the nucleic acid is delivered is a human cell.

15. A method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule (RNAi), said anti-sense molecule, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for said protein, wherein absent said anti-sense molecule, ribozyme or RNAi nucleic acid, the levels of said protein or its mRNA is less than in a non-Peyer's patch cell.

16. The method of Claim 15 wherein the anti-sense nucleic acid, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule is complementary to a sequence of at least 15 nucleotides of the mRNA of the protein.

17. The method of Claim 16 wherein the anti-sense nucleic acid, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule is complementary to a sequence of at least 30 nucleotides of the mRNA of the protein.

18. The method of Claim 15 wherein the protein is coded for by a gene with an expression Fold Change denoted by a "-", or a number less than 0.5 in Tables 2 or 3.

19. The method of Claim 15 comprising delivering to said cell anti-sense nucleic acid molecules, ribozyme nucleic acid molecules, RNA interference nucleic acid molecules, said anti-sense, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for at least 5 different protein a, wherein absent said anti-sense, ribozyme or RNAi nucleic acid molecule, the levels of each of said proteins or its mRNA is less than in a non-Peyer's patch cell.

20. A method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule said anti-sense, ribozyme or RNAi nucleic acid forming a double-stranded molecule with part or all of the mRNA for said protein, wherein absent said anti-sense, ribozyme or RNAi nucleic acid molecule, the levels of said protein or its mRNA is less than in a non-Peyer's patch cell.

21. A method of Claims 1, 13, or 15 in which the Peyer's patch cell is an M cell.

22. A human cell to which the method of Claims 1 has been applied, or the progeny of said human cell.

23. A human cell to which the method of Claim 13 has been applied, or the progeny of said human cell.

24. A human cell to which the method of Claim 15 has been applied, or the

progeny of said human cell.

25. A human cell to which the method of Claims 1 has been applied, or the progeny of said human cell.

5 26. A human cell to which the method of Claim 13 has been applied, or the progeny of said human cell.

27. A human cell to which the method of Claim 15 has been applied, or the progeny of said human cell.

10 28. A method for enhancing transport of a drug through the gastrointestinal tract, said method comprising orally administering said drug in a composition that comprises a transport-enhancing protein, said transport-enhancing protein selected from the group consisting of human serum albumin (HSA), clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, and Ca²⁺pla2, or a homolog that has at least 80% amino acid identity with said transport-enhancing protein over a length of said transport-enhancing protein identical to the homolog.

15 29. A method of Claim 28 wherein the homolog has at least 90% amino acid with the transport-enhancing protein over a length of the transport-enhancing protein identical to the homolog.

20 30. A method of Claim 28 wherein the transport-enhancing protein is selected from the group consisting of human serum albumin (HSA), clusterin, T-cell surface glycoprotein CD5 precursor, HSP84, and Ca²⁺pla2.

31. A method to facilitate intracellular trafficking of an antigen that has been orally delivered by itself or as part of a composition or particle, said method comprising administering a protein selected from the group consisting of calreticulin, rab family proteins and ribosomal proteins.

25 32. A chimeric protein comprising the amino acid sequence for calreticulin, rab family proteins and ribosomal proteins and the amino acid sequence for a second polypeptide.

33. A method of administering a polypeptide, where said polypeptide is part of a chimeric protein of Claim 32, and wherein said chimeric protein is orally administered.

30 34. A method of delivering a vaccine to a target cell, said method comprising utilizing as the target cell a Peyer's patch cell in which a normally upregulated protein or mRNA is further upregulated.

35. A method of Claim 34 wherein the Peyer's patch cell is an M Cell.

36. A method of Claim 1 wherein the protein is selected from the group consisting

of clusterin, T-cell surface glycoprotein CD5 precursor, HSP 84, Ca²⁺ dependent phospholipase A2 precursor, and the ribosomal proteins, S12, S11, L12, L11, S29, S19, L21, L19, L13, L44, and L36.

37. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of clusterin, T-cell surface glycoprotein CD5 precursor, HSP 84, and Ca²⁺ dependent phospholipase A2 precursor and the mRNA is for a protein selected from said group.

38. A method of Claim 1 wherein the protein is selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, CD40, C-MYC, PKC- α , GSTA1, GATA-2, PLGF, ezrin, HGF activator, hepatocyte growth factor-like protein, NCAD, MNDA, LHX1, TIE-1, NCAML1, CD104, CD44, SRC1, NMMA, TKT, ephrin (type A), Sp1, RAB proteins, PKC, TIR, Jak3, EGR-1, TNK1, CAMK IV, HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd153, XPG, XPD, ubiquitin-conjugating enzyme, RAD23, cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, α 3 β 1 integrin, CD11C antigen, CD104 antigen, CD44, NMMA, TKT, ephrin (type A), and Sp1, a RAB protein, PKC, TfR, bcl-x and caspase-9.

39. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, CD40, C-MYC, PKC- α , GSTA1, GATA-2, PLGF, ezrin, HGF activator, hepatocyte growth factor-like protein, NCAD, MNDA, LHX1, TIE-1, NCAML1, CD104, CD44, SRC1, NMMA, TKT, ephrin (type A), Sp1, RAB proteins, PKC, TIR, Jak3, EGR-1, TNK1, CAMK IV, HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd153, XPG, XPD, ubiquitin-conjugating enzyme, RAD23, cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, α 3 β 1 integrin, CD11C antigen, CD104 antigen, CD44, NMMA, TKT, ephrin (type A), and Sp1, a RAB protein, PKC, TfR, bcl-x and caspase-9. and the mRNA is for a protein selected from said group.

40. A method of Claim 1 wherein the protein is selected from the group consisting of an IL-2 receptor, a gamma c chain of an IL-2 receptor, interferon - γ , and a C-C chemokine.

41. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of an IL-2 receptor, a gamma c chain of an IL-2 receptor, interferon - γ , and a C-C chemokine and the mRNA is for a protein selected from said group.

42. A method of Claim 1 wherein the protein is selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, PKC- α , GSTA1, GATA-2, and PLGF.
43. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of cyclin D1, PLC-L, GRB2, ERK3/MAPK6, ERK1, ERK3, JNK2, CD40, CRAF1, C-MYC, PT- α , IL-R, PKC- α , GSTA1, GATA-2, and PLGF and the mRNA is for a protein selected from said group.
44. A method of Claim 1 wherein the protein is selected from the group consisting of a RAB protein, PKC, and TfR.
45. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of a RAB protein, PKC, and TfR and the mRNA is for a protein selected from said group.
46. A method of Claim 1 wherein the protein is selected from the group consisting of Jak 3, EGR-1, TNK1, and CAMK IV.
47. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of Jak 3, EGR-1, TNK1, and CAMK IV and the mRNA is for a protein selected from said group.
48. A method of Claim 1 wherein the protein is selected from the group consisting of HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd 153, XPG, XPD, ubiquitin, conjugating enzyme, RAD 23, and ataxia telangiectasia.
49. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of HSP40, HSP70, HSP60, HSO27, fMLP-related receptor, HSP27, glutaredoxin, CREB, gadd 153, XPG, XPD, ubiquitin, conjugating enzyme, RAD 23, and ataxia telangiectasia and the mRNA is for a protein selected from said group.
50. A method of decreasing the levels of a protein in a Peyer's patch cell, said method comprising delivering to said cell a DNA molecule coding for an anti-sense nucleic acid molecule, a ribozyme nucleic acid molecule, an RNA interference nucleic acid molecule (RNAi), said anti-sense molecule, ribozyme or RNAi nucleic acid being complementary to a sequence of at least 10 nucleotides of the mRNA for said protein, wherein absent said anti-sense molecule, ribozyme or RNAi nucleic acid, the levels of said protein or its mRNA is less than in a non-Peyer's patch cell.
51. A method of increasing the extent to which the function of a protein is carried out in a Peyer's patch cell, said method comprising delivering to said cell a nucleic

acid coding for said protein, wherein absent said delivery, the level of said protein or its mRNA is greater in said cell than in a non-Peyer's patch cell.

52. A chimeric protein that comprises two or more segments, each of said segments enhancing a different step in the peptide transport process, said steps selected from the group consisting of binding to a cell, transporting the peptide into the cell, transporting the peptide through the cell, and transporting the peptide out of the cell.

53. A chimeric protein of Claim 52 wherein one of the segments binds to the cell.

54. A chimeric protein of Claim 52 wherein one of the segments is a protein that is more prevalent in a Peyer's patch cell than in a non-Peyer's patch cell.

55. A chimeric protein of Claim 52 wherein the cell is a Peyer's patch cell.

56. A chimeric protein of Claim 55 wherein the cell is an M cell.

57. A method of targeting a composition or delivery vehicle to a Peyer's patch cell said method comprising utilizing a composition or vehicle that contains a protein ligand that will specifically bind to a protein that is up-regulated in Peyer's patch cells.

58. The method of Claim 57 wherein the composition or delivery vehicle comprises a drug or antigen.

59. A method of selecting for a ligand that will selectively bind to a target in a Peyer's patch cell, said method comprising contacting a phage library with a protein that is upregulated in Peyer's patch cells.

60. The method of Claim 59 wherein the protein is attached to a solid substrate.

61. A method of Claim 1 wherein the protein is selected from the group consisting of HGF activator, ezrin, NCAD, MNDA, and LHX1.

62. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of HGF activator, ezrin, NCAD, MNDA, and LHX1, and the mRNA is for a protein selected from said group.

63. A method of Claim 1 wherein the protein is selected from the group consisting of cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, $\alpha 3\beta 1$ integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1.

64. A method of Claim 34 wherein the upregulated protein is selected from the group consisting of cadherin 2, neural cell adhesion molecule, integrin alpha 3, leukocyte adhesion glycoprotein p150, integrin beta 4, TIE, NCAML1, $\alpha 3\beta 1$ integrin, CD11C antigen, CD104 antigen, CD44, NMDA, TKT, ephrin (type A), and Sp1, and the mRNA is for a protein

selected from said group.

- 5 65. A method of promoting enterocyte-M cell conversion, said method comprising orally administering an antigen, antigenic composition, or antigen-carrying particle to a person and either simultaneously with, or prior to, said administration, also orally administering a bacteria, or pro-biotic yogurts, or bacterial component to said person.

AMINO ACID SEQUENCES AND NUCLEOTIDE SEQUENCES CORRESPONDING TO SELECTED
GENBANK ID NUMBERS

5

GENBANK ID: M81750
VERSION M81750.1 GI:895928

10

MVNEYKKILLLLKGFELMDDYHFTSIKSLLAYDLGLTTKMQEEYN
RIKITDLMEKKFQGVACLOKLIELAKDMPSLKNLVNNLRKEKSKVAKKIKTQEKAPVK
KINQEEVGLAAPAPTARNKLTSEARGRIPVAQKRKTPNKEKTEAKRNKVSQEQSKPPG
PSGASTSAVDHPPLPQTSSSTPSNTSFTPNQETQAQRQVDARRNVPQNDPVTVVVLK
ATAPFKYESPENGKSTMFHATVASKTQYFHVKVFDINLKEKFVRKKVITISDYSECKG
15 VMEIKEASSVSDFNQNFVFNRIEIANKTPKISQLYKQASGTMVYGLFMLQKKSVMK
KNTIYEIQDNTGSMDDVVGSGKWHNIKCEKGDKLRLFLCLQLRTVDRKLKLVCGSHSFIK
VIKAKKNKEGPMNVN

20

GENBANK ID: X59798
VERSION X59798.1 GI:35631

25

MEHQLLCCEVETIRRAYPDANLLNDRVLRAMLKAEETCAPSVSY
FKCVQKEVLPMSRKIVATWMLLEVCEEQKCEEEVFPLAMNYLDRFLSLEPVKKSRLQLL
GATCMFVASKMKETIPLTAEKLCIYTDNSIRPEELLQMEILLVNLKWNLAAMTPHDF
IEHFLSKMPEAEENKQIRKHAQTFVALCATDVKFISNPPSMVAAGSVVAAVQGLNLR
SPNNFLSYRLTRFLSRVICKDPCDLRACQEQIEALLESSLRQAQQNMDPKAAEEEE
EEEEVDLACTPTDVRDVI

30

GENBANK ID: L27211.1
VERSION L27211.1 GI:558656

35

MEPAAGSSMEPSADWLATAAARGRVEEVRLLEAGALPNAPNSY
GRRPIQVMMGSARVAELLLHGAEPNCADPATLTRPVHDAAREGFDTLVVLHRAGA
RLDVRDAWGRLPVDLAEELGHRDVARYLRAAAGGTRGSNHARIDAAEGPSDIPD

45

GENBANK ID: U22398
VERSION U22398.1 GI:790247

40

MSDASLRSTSTMERLVARGTFPVLVRTSACRSLEFGPDHEELSR
ELQARLAELNAEDQNRWDYDFQQDMPLRGPGRLQWTEVDSDSVPAFYRETVQVGRCL
LLAPRPVAVAVAVSPPLEPAAESLDGLEEAPEQLPSVPVPAPASTPPVPVLPAPAPAP
APAPVAAPVAAPVAVAVLAPAPAPAPAPAPAPAPVAAPAPAPAPAPAPAPAPAPDA
APQESAEQGANQGQEQEPLADQLHSGISGRPAAGTAAASANGAAIKKLSGPLISDFF
AKRKRSAPEKSSGDVPAPCPSPSAAPGVGSVEQTPRKRLR

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GENBANK ID: X51521
VERSION X51521.1 GI:31282

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MPKPINVRVTMDAELEFAIQPNTTGKQLFDQVVKITIGLREVWY
FGLHYVDNKGFPWLKLDKKVSAQEVVRKENPLQFKFRAKFYPEDVAEELIQDITQKLF
FLQVKEGILSDEIYCPPETAVLLGSYAVQAKFGDYNKEVHKSGYLSSERLIPQVMDQ
HKLTRDQWEDRIQVWHAHRGMLKDNAMLEYLKIAQDLEMYGINYFEIKNKKGTDLWL
GVDALGLNIYEKDDKLTPIKIGFPWSEIRNISFNDKKFVIKPIDKKAPDFVFYAPRLRI
NKRIQLQCMGNHELYMRRRKPDITIEVQQMKAQAREEKHQKQLERQQLETEKKRRRETVE
REKEQMREKEELMLRLQDYEEKTKAERELSEIQRALQLEERKRAQEEAERLEAD
RMAALRAKEELERQAVDQIKSQEQLAELAETAKIALLEEARRRKEDEVEEWQHRAK
EAQDDLVTKEELHLVMTAPPPPPPPVYEPVSYHVQESLQDEGAEPYGYSAELSSEGI
RDDRNEEKRITEAEKNERVQRQLVTLSSSELSQARDENKRTHNDIIHNENMRQGRDKYK
TLRQIRQGNTKQRIDEFEAL

60

GENBANK ID: L04143.1
VERSION L04143.1 GI:180574

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THIS ENTRY IS NOT CONTIGUOUS GENOMIC DNA. IT CONTAINS NUMEROUS PIECES OF
INTRONS.

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55 GENBANK ID: M32110
VERSION M32110.1 GI:189421

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5 RFRERRFHPSLRSTRRFYPHTHNMDGFFIAKFKKFSNSIQSQTGNSETATPTNVDLP
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10 GenBank ID: J04111
VERSION J04111.1 GI:186624

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20 GENBANK ID: X59932
VERSION X59932.1 GI:30255

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30 GENBANK ID: L29220
VERSION L29220.1 GI:632969

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40 GENBANK ID: M91815.1
VERSION M91815.1 GI:180169
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61 CAAGGATGGA TGAAACTGGA AAAGAATGAA AGGACCCCTT ATATCATGAA AACCACTAAG
121 CACTTCAATG ACATCAGTAA CTTGATTGCT TCAGAAATCA TCCGCAATGA GGACATCAAC
181 GCCAGGGTGA GCGCCATCGG GAAGTGGGTG GCCGTAGCTG ACATATGCCG CTGCCTCCAC
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50 541 ATCCGAGAGA TTCGCCAGTT TCAACAACT GCCTACAAA TAGAGCACCA AGCAAAGGTA
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55 GENBANK ID: M26708
VERSION M26708.1 GI:190695

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65 GENBANK ID: M81757
VERSION M81757.1 GI:337732

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5 GENBANK ID: V00568
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 VERSION V00568.1 GI:34815

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GENBANK ID: NM_011587.1
VERSION NM_011587.1 GI:6755784

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MVWWGSSLLPTLFLASHVGASVDLTLLANLRITDPQRFELTCV
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SYENTLDWQEADDGRFQLQLQNVQPPSSGIYSATYLEASPLGSAFFRLIVRGCGAGRW
GPGCVKDCPGCLHGGVCHDHDGECVCPPGFTGTRCEQACREGRFQSCQEQCPGTAGC
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SGWHGVHCEKSDRIPQILSMATEVEFNIGTMPRINCAAAGNFPVRGSMKLRKPDGTM
LLSTKVIVEPDRTTAEFEVPSLTLDGDSGFWECEVSTSGGQDSRRFKVNVKVPVPLTA
PRLAKQSRQLVVSPLVSFSGDGPISSVRLHYRPQDSTIAWSAIVVDPSENVTLMLNK
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DPVRESRAAEEGLDQQLVLAVVGSVSATCLTILAALLALVCIRRSCLHRRRTFTYQSG
SGEETILQFSSGTLTLTRPKPQPEPLSYPVLEWEDITFEDLIGEGNFGQVIRAMIKK
DGLKMNAAIKMLKEYASENDRDFAGELEVLCCKLGHHPNIINLLGACENRGYLYIAIE
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HRDLAARNVLVGENLASKIADFGLSRGEEVYVKKTMGRLPVRWMAIESLNYSVYTTKS
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RPYERPPFAQIALQLGRMLEARKAYVNMSLFENFTYAGIDATAEEA

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GENBANK ID: NM_002867.1
VERSION NM_002867.1 GI:4506368

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MASVTDGKHGVKDASDQNFDMFKLLIIGNSSVGKTSFLLRYAD
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GENBANK ID: P27361
NO VERSION DATA

5 LNENQKLAVKRILSGDCRPLPYILFGPPGTGKTVTIIIEAVLQVH
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GFPLIFHGVRGSEAREGKSPSWFNPAEAVQVLRVYCCLLAHSISSQVSASDIGVITPYR
KQVEKIRILLRNVDLMDIKVGSVEEFQGEYLVIISTVRSNEDRFEDDRYFLGFLSN
SKRFNVAITRPKALLIVLGNPHVLVRDPCFGALLEYSITNGVYMGCDLPPALQSLQNC
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10 GENBANK ID: NM_002752.1
VERSION NM_002752.1 GI:4506096

15 MSDSKCDSQFYSVQVADSTFTVLKRYQQLKPIGSGAQGIVCAAF
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DVYLMELMDANLCQVIHMELDHERMSYLLYQMLCGIKHLHSAGIIHRDLKPSNIVVK
SDCTLKILDFGLARTACTNFMPTPYVVTRYRAPEVILGMGYKENVDIWSVGCIMGEL
VKGCVIFQGTDHIDQWNKVIEQLGTPSAEFMKKLOPTVRNYVENRKPYPGKFEELFP
DWIFPSESERDKIKTSQARDLLSKMLVIDPDKRISVDEALRHPYITVWYDPAEAEAPP
20 PQIYDAQLEEREHAIEEWKELIYKEVMDWEERSKNGVVKDQPSAQMQQ

25 GENBANK ID: M22382.1
VERSION M22382.1 GI:190126

30 MLRLPTVFRQMRPVSRVLAPHLTRAYAKDVKFGADARALMLQGV
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PVTTPPEIAQVATISANGDKEIGNIISDAMKKVGRKGVITVKDGKTLNDELEIIEGMK
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35 DVDGEALSTLVNRLKVLQVAVKAPFGDNRNKQLKDMAIATGGAVFGEGLTLNL
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40 GENBANK ID: U09564.1
DEFINITION HUMAN SERINE KINASE MRNA, COMPLETE CDS.
VERSION U09564.1 GI:507212

45 MERKVLALQARKKRTKAKKDKAQRKSETQHRGSAPHSESIDLPEQ
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50 TEDIQTRQYRSLEVLIGSGYNTPADIWSTACMAFELATGDYLFEPHSGEYTRDEDHI
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55 GENBANK ID: M11507.1
VERSION M11507.1 GI:339515

60 MMDQARSFAFSNLFGGEPLSYTRFSLARQVDGDNHSHVEMKLAVDE
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GTESPVREEPGEDFPAARRLYWDDLKRKLSEKLDSTDTSTIKLLNENSYPREAGSQ
KDENLALYVENQFREFKLSKVWRDQHFVKIQVKDSAQNSVIIVDKNGRLVYLVENPGG
YVAYSKAATVTGKLVHANFGTKKDFEDLYTPVNGSIVIVRAGKITFAEKVANAESLNA
IGVLIYMDQTKFPIVNAELSFFGHAHLGTDPYTPGFPSFNHTQFPSSRSGLPNIPV
QTISRAAAEKLFGNMEGDCPSDWDKTDSTCRMVTSSEKNVKLTVSNVLKEIKILNIFGV
IKGFVEPDHYVVVGAQRDAWGPGAAGSGVGTALLKLAQMFSDMVLKDGFPQSRSTIF
65 ASWSAGDFGSGATEWLEGYLSLHLKAFTYINLDKAVLGTSNFKVSASPLLYTLIEK
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GTTMDTYKELIERIPELNKVARAAAEVAGQFVIKLTHTDVELNLDYERYNSQLLSFVRD
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YHFLSPYVSPKESPFHRVFWGSGSHTLPALLENLKLKQNNGAFNETLFRNQLALATW
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GENBANK ID: U55017.1
VERSION U55017.1 GI:1297296

MESYHKPDQKQLQALKDTANRLRISSIQATTAAGSGHPTSCCSA
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ISSDLGHPVPKQAFDVAATGSLGQGLGAACGMAYTGKYFDKASYRVYCLLDGELSE
GSVWEAMAFASIYKLDNLVAILDINRLGQSDPAPLQHQMIDIYQKRCEAFGWHAIIIDG
HSVEELCKAFGQAKHQPTAIIAKTFKGRGITGVEDKESWHGKPLPKNMAEQIIQEIYS
QIQSKKILATPPQEDAPSVDIANIRMPSLPSYKVGDKIATRKAYGQALAKLGHASDR
IIALDGDTKNSTFSEIFKKEHPDRFIECYIAEQNMVSIAGCATRNRTVPFCSTFAAF
FTRAFDQIRMAAISESNINLCGSHCGVSIAGEDGPSQMALEDLAMFRSVPTSTVFYPSD
GVATEKAVELAANTKGICFIRTSRPENAIYNNNEDFQVGQAKVVLKSKDDQVTVIGA
GVTLHEALAAELLKKEKINIRVLDPFTIKPLDRKLILDSARATKGRILTVEDHYEG
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GENBANK ID: X14034.
VERSION X14034.1 GI:35513

MSTTVNVDSLAEYEKSQIKRALELGTVMTVFSFRKSTPERRTVQ
VIMETRQVAWSKTADKIEGFLDIMEIKEIRPGKNSKDFERAKAVRQKEDCCFTILYGT
QFVLSTLSLAADSKEDAVNWLSSLKILHQAAMNASTPTIIESWLRKQIYSVDQTRNS
ISLRELKTIPLINFKVSSAKFLKDKFVEIGAHKDELSFEQFHLFYKKLMFEQQKSIL
DEFKKDSSVFILGNTDRPDASAVYLHDFQRFLIHEQQEHWAQDLNKVRERMTKFIDDT
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QLRSESSPEAYIRCLRMGCRCIELDCWDGPDGKPVIIHGWTTRTKIKFDDVVAIKDH
AFVTSSFPVILSIEEHCSVEQQRHMAKAFKEVFGDLLLTKPTEASADQLSPSPQLREK
IIIKHKKLGPRGDVDVNMEDKKDEHKQOGELYMWDSIDQKWRHYCAIADAKLSFSDD
IEQTMEEVFPQDIPTELHFGEKWFHKKVEKRTSAEKLQEYCMETGGKDGTFVRES
ETFPNDYTLSEWRSRGRVQHCRIRSTMEGGTLKYLYLDNLRFRMYALIQHYRETHLPC
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ERYNTERDINSLYDVSVMYVDPSEINPSMPQRTVKALYDYKAKRSDELSFCRGALIH
VSKEPGGWWKGDYGTIRIQYFSPNYVEDISTADFEELKQIIEDNPLGSLCRGILDIN
TYNVVKAPOGKNQKSFVFILEPKEQGDPEFATDRVEELFEWFQSIREITWKIDSKE
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NGRTGYVLQPEMRTEKYDPMPPESQRKILMTLTVKVLGARHLPKLGRSIACPFVEVE
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FLAHATYPIKAVKSGFRSVPLKNGYSEDIELASLLVFCEMRPVLESEELYSSCRQLR
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GENBANK ID: M27691.1
VERSION M27691.1 GI:181038

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YQTSSGQYIAITQGGAIQLANNGTDGVQGLQTLTMTNAAATQPGTTILQYAQTTDGQQ
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GENBANK ID: M18391
VERSION M18391.1 GI:339716

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PGPAGLVEVAGTCLPHARASPRPSGAPRMHCSPDGEWLVPVGRCHCEPGYEEGGSGEA
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5 SPGARALTTPAVHVNGLEPYANYTFNVEAONGVSGLGSSGHASTSVSISMGHAESLSG
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RELDPAWLMVDTVIGEGEFGEVYRGTLRLPSQDCKTVAIKTLKDOTSPGGQWWNFLREA
TIMGQFSHPHILHLEGVVTKRKPIIMIITEFMENAALDAFLREREDQLVPGQLVAMLOQ
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10 RWTAPETIAHRIFFTASDVWSFGIVMWEVLSFGDKPYGEMSNQEVMSIEDGYRLPPP
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15 GENBANK ID: X54079.1
VERSION X54079.1 GI:32477

MTERRVPFSLLRGPSWDPFRDWYPHSRLFDQAFGLPRLPEEWSQ
WLGGSSWPGYVRPLPPAAIESPAVAAPAYSRLSRQLSSGVSEIRHTADRWRVSLDVN
HFAPDELTVKTKDGVVEITGKHEERQDEHGYSRCFTRKYTLPPGVDPQTQVSSLSPE
20 GTLTVEAPMPKLATQSNEITIPVTFESRAQLGGPEAAKSDETAAK

GENBANK ID: XM_012654.3
VERSION XM_012654.3 GI:14773503

25 MFGVTLWEMFSGGEEPWAGVPPYLILQRLDRARLPRPPLCSRA
LYSLALRCWAPHSRPSFSHLEGLLQEAGPSEACVRDVTEPGALRMETGDPITVIE
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30 GENBANK ID: XM_054457.2
VERSION XM_054457.2 GI:18590931

35 MASATDSRYGQKESSDQNFDMFKILIIGNSSVGKTSFLFRYAD
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40 GENBANK ID: XM_038595.3
VERSION XM_038595.3 GI:18590923

45 MAPPSEETPLIPQRSCSLSTEAGALHVLLPARGPGPPQRLSFS
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55 RQQLPAPKWTELALLIQCMAYEPVQRPSFRAVIRDLSLISSDYELLSDPFGALAP
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QRHRARLDASRLLYSSQICKGMEYLGSRRCVHRDLAARNILVESEAHVKIADFGLAK
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60 SAEFLRMMGCERDVPALCRLLLEELLEGQRLPAPPACPAEVHELMKLCWAPSPQDRPSF
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65 GENBANK ID: NM_002755.2
VERSION NM_002755.2 GI:14589898

5 MPKKKPTPIQLNPAPDGSVNGTSSAETNLEALQKKLEELDE
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10 GENBANK ID: NM_001744.1
VERSION NM_001744.1 GI:4502556

15 MLKVTVPSCSASSCSSVTASAAPGTASLVPDYWIDGSNRDALS
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GPEVDMWSVGIIITYILLCGFEPFYDERGDQFMFRRILNCEYFISPWDEVSLNAKDL
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25 GENBANK ID: XM_053461.2
VERSION XM_053461.2 GI:18553657

30 MASRGATRPNGPNTGNKICQFKLVLLGESAVGKSSLVLRVFKGQ
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35 GENBANK ID: U33635.1
VERSION U33635.1 GI:1016701

40 MGAARGSPARPRRLPLLSVLLPLLLGGTQTAIVFIKQPSSQDAL
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IADESFAVVLAPQDVVVARYEAMFHCQFSAQPPPSLQWLFEDETPITNRSRPPHLR
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45 AGSEERVTCCLPPKGLPEPSVWWEHAGVRLPTHGRVYQKGHELVLANIAESDAGVYTCH
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WKGKDRILDPTKLGPRMHIFQNGSLVIHVDVAPEDSGRYTCIAGNSCNKIKHTEAPLYV
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50 NVVRLGLCREAPHYMVLEYVDLEDLQFLRISKSKDEKLKSQPLSTKQKVALCTQV
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55 GENBANK ID: XM_004559.5
VERSION XM_004559.5 GI:17464405

60 MGPEALSSLLLLLLVASGDADMKGHFDPAKCRYALGMQDRTIPD
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65 ERRVLEELTVHLSVPGDTILINNRPGPREPPYQEPNPRGNPPHSAPCVPNGSALLL
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5 PHYAEADIVTLQGVTTGGNTYAVPALPPGAVGDGPPRVDFPRSRRLRFKEKLGEQGFGEV
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10 GENBANK ID: M28212.1
VERSION M28212.1 GI:550071

/GENE="RAB6"

CDS 71..697

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15 1 AGCTGGCTGG AGCAGCATCG GTCCGGGACG GTCTCTAGGC TGAGGCGGCG GCCGCTCCTC
61 TAGTTCCACA ATGTCCACGG GCGGAGACTT CGGGAATCCG CTGAGGAAAT TCAAGCTGGT
121 GTTCCTGGGG GAGCAAAGCG TTGGAAAGAC ATCTTTGATC ACCAGATTCA TGTATGACAG
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50 7261 AATGCAGCAG CTGAAGGGCT TTGTGGTGAG GGCAATGACC CTTGAAGATA TTCAGACCAG
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60 7921 CAGATCTTAG GGATGATTAA AGGCAGCATT TGATGATAGC AGACATTGTT ACAAGGACAT
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10 GENBANK ID: S40706.1
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20 GENBANK ID: M96995.1
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35 GENBANK ID: U32944.1
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40 GENBANK ID: NM_021141.2
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60 GENBANK ID: NM_005053.1
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5 GENBANK ID: Z23115.1
VERSION Z23115.1 GI:510900

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10 GENBANK ID: AB020979.1
VERSION AB020979.1 GI:6518501

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20 GENBANK ID: U21092.1
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35 GENBANK ID: NM_001459.1
VERSION NM_001459.1 GI:4503750

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45 GENBANK ID: X57500.1
VERSION M36089.1 GI:340396

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60 GENBANK ID: BAA02962.1
DEFINITION HUMAN MRNA FOR RECA-LIKE PROTEIN HSRAD51, COMPLETE CDS.

VERSION D13804.1 GI:397826
CDS 212..1231
35 /CODON_START=1

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30 GENBANK ID: B56529

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VERSION X84740.1 GI:860962
45 CDS 334..3102
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VERSION M62829.1 GI:182262

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GENBANK ID: U10421.1
VERSION U10421.1 GI:500756

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GENBANK ID: U08015.1
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GENBANK ID: M55654
VERSION M55654.1 GI:339491

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GENBANK ID: NM_005568.1
VERSION NM_005568.1 GI:5031866

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GENBANK ID: X69111.1
VERSION X69111.1 GI:32294

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GENBANK ID: NP_000507.1
VERSION NP_000507.1 GI:4504047

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181 KIDVIKQADY VPSQDQLLRC RVLTSIGIFET KFQVDKVNFB MFDVGGQRDE RRKWIQCFND
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GENBANK ID: AAA40889.1
VERSION AAA40889.1 GI:203357

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GENBANK ID: B53771
VERSION B53771 GI:2136296

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GENBANK ID: M92299.1
VERSION M92299.1 GI:184292

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GENBANK ID: M68891.1
VERSION M68891.1 GI:182995

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GENBANK ID: XM_028606.2
VERSION XM_028606.2 GI:15304625

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GENBANK ID: NP_009077.1
DEFINITION HOMO SAPIENS ZINC FINGER PROTEIN 161 (ZNF161), MRNA.
VERSION NM_007146.1 GI:6005967
CDS 42..1592
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541 GTAAGCCTGT CAAGAAGAAC CATGCTTGTG AGATGTGTGG GAAGGCCTTC CGAGATGTGT
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661 GTAATCAGCG CTTCAAGAGG AAGGACCGGA TGACTTACCA TGTGAGGTCT CATGAAGGAG
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841 CTGCTGCCTT TGCCACCAA GACAGACTGC GGACACACAT GGTGCGCCAT GAAGGCAAGG
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GENBANK ID: AAA36598.1
DEFINITION HUMAN STEM CELL PROTEIN (SCL) MRNA, COMPLETE CDS.
VERSION M29038.1 GI:337958
CDS 81..725
/CODON_START=1

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35 GENBANK ID: NP_006168
DEFINITION HOMO SAPIENS NEURAL RETINA LEUCINE ZIPPER (NRL), MRNA.
VERSION NM_006177.1 GI:5453801
CDS 118..831
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GENBANK ID: AAA58399.1
VERSION M95809.1 GI:179568
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CDS 55..1701
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55 GENBANK ID: AAA61146.1
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20 GENBANK ID: M27492.1
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35 GENBANK ID: L34059
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GENBANK ID: M81695.1

VERSION M81695.1 GI:487829

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GENBANK ID: X51841.1

VERSION X51841.1 GI:33910

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VERSION XP_030326.1 GI:14763626

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301 EDFISSTIST TPRAFDHTKQ NQDWTQWNPS HSNPEVLLQT TTRMTDVRN GTTAYEGNWN
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GENBANK ID: NP_000826.1
VERSION NP_000826.1 GI:4504129

1 MGGALGPALL LTSLEFGAWAG LGPGQGEQGM TVAVVFSSSG PPQAQFRARL TPQSFLDLPL
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601 KSVWLLWALV FNNSVPIENP RGTTSKIMVL WVAFFAVIFL ASYTANLAAF MIQEYIDTV
661 SGLSDKKFOR PQDQYPPFR GTVPNGSTER NIRSNDYDMH THMVKNQORS VEDALTSKLM
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841 WKLRHVPNS SOLDFLLAFS RGIYSCFSGV QSLASPPRQA SPDLTASSAQ ASVLKMLQAA
901 RDMVTTAGVS SSLDRATRTI ENWGGGRRAP PPSPCPTPS GPSPCLPTPD PPPEPSPTGW
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1141 AGAPAWQHRQ HVCLHAHAHL PFCWGAVCPH LPPCASHGSW LSGAWGPLGH RGRTLGLGTG
1201 YRDSGGLDEI SSVARGTQGF PGPCTWRRIS SLESEV

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GENBANK ID: CAA43045.1

DEFINITION HUMAN CDW40 MRNA FOR NERVE GROWTH FACTOR RECEPTOR-RELATED
B-LYMPHOCYTE ACTIVATION MOLECULE.

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VERSION X60592.1 GI:29850

CDS 48..881

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781 CCAACACTGC TGCTCCAGTG CAGGAGACTT TACATGGATG CCAACCGGTC ACCCAGGAGG
841 ATGGCAAAGA GAGTCGCATC TCAGTGACAG AGAGACAGTG AGGCTGCACC CACCCAGGAG
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GENBANK ID: AAB59544.1

DEFINITION HUMAN NERVE GROWTH FACTOR RECEPTOR MRNA, COMPLETE CDS.

VERSION M14764.1 GI:189204

CDS 114..1397

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601 CGTGCCCTGCC CTGCACCGTG TGCGAGGACA CCGAGCGCCA GCTCCGCGAG TGCACACGCT
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901 TTGTGGGCCT TGTGGCCTAC ATAGCCTTCA AGAGGTGGAA CAGCTGCAAG CAGAACAAGC
961 AAGGAGCCAA CAGCCGGCCA GTGAACCAGA CGCCCCACC AGAGGGAGAA AAACCTCCACA
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2221 GCTTGCCCTAG GGCCTGGTCC ATGATGGAGT CAGGTTTGGG GTTCGTGGAA AGGGTGCTGC
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GENBANK ID: Q00941
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10 GENBANK ID: P05106
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CCACACTCAAAGGTCCCATTGCCATTGTTGCAGCGATGGCTATTATGTTGAGCTTGGGCC
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55 GGCTTTATGGTAAAGGAATTTCTCCTTTCTTGGGGACAGACTCGCACCTTGGCCCTAATG
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60 GENBANK ID: CAA52348.1
VERSION X74295.1 GI:437781
CDS <1..234
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35 1 AAGATGGGAT TCTTCAAACG GGCGAAGCAC CCCGAGGCCA CCGTGCCCCA GTACCATGCG
61 GTGAAGATTC CTCGGGAAGA CCGACAGCAG TTCAAGGAGG AGAAGACGGG CACCATCCTG

5
121 AGGAACAACCT GGGGCAGCCC CCGGCGGGAG GGCCCGGATG CACACCCCAT CCTGGCTGCT
181 GACGGGCATC CCGAGCTGGG CCCCAGTGGG CATCCAGGGC CAGGCACCGC CTAGGTTCCC
241 ATGTCCCAGC CTGCGCTGTG GCTGCCCTCC ATCCCTTCCC CAGAGATGGC TCCTTGGGAT
301 GAAGAGGGTA GAGTGGGCTG CTGGTGTAC ATCAAGAATT TGGCAGGATC GGCTTCCTCA
361 GGGGCACAGA CCTCTCCAC CCACAAGAAC TCCTCCACC CAACTTCCC TTAGAGTGCT
421 GTGAGATGAG AGTGGGTAAA TCAGGGACAG GGCCATGGGG TAGGGTGAGA AGGGCAGGGG
481 TGTCTGATG CAAAGGTGGG GAGAAGGATC CTAATCCCTT CCTCTCCCAT TCACCTGTG
541 TAACAGGACC CCAAGGACCT GCCTCCCCG AAGTGCCTTA ACCTAGAGGG TCGGGGAGGA
601 GGTGTGTCA CTGACTCAAG GCTGCTCCTT CTCTAGTTTC CCCTCTCATC TGACCTTAGT
10 661 TTGCTGCCAT CAGTCTAGT GTTTCGTGGT TTCGTCTATT TATTAAAAA TCGGAACCC

GENBANK ID: M57627
VERSION CAA51942.1 GI:580177
1 MHSSAL

15
GENBANK ID: AAA52578.1
VERSION AAA52578.1 GI:183364

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1 MWLQSLLLLG TVAC SISAPA RSPSPSTQPW EHVNAIQEAR RLLNLSRDTA AEMNETVEVI
61 SEMFDLQEP CLQTRLELYK QGLRGLSLTKL KGPLTMMASH YKQHCPTPE TSCATQIITF
121 ESFKENLKDF LLVIPFDCWE PVQE

25
GENBANK ID: AAA58482.1
VERSION AAA58482.1 GI:182669

30
1 METNFSIPLN ETEEVLPEPA GHTVLWIFSL LVHGVTFVFG VLGNGLVIWV AGFRMTRTVN
61 TICYLNLA DEFESAILPF RMVSVAMREK WPFASFCKL VHVIMIDINLF VSVYLITIIA
121 LDRCICVLHP AWAQNHRTMS LAKRVTGLW IFTIVLTLPN FIFWTTIST NGDYCIFNF
181 AFWGDTAVER LNVFITMAKV FLILHFIIGF TVPMSIITVC YGIIAAKIHR NHMIKSSRPL
241 RVFAAVVASF FICWFYELI GILMAVWLKE MLLNGKYKII LVLINPTSSL AFFNSCLNPI
301 LYVFMGRNFQ ERLIRSLPTS LERALTEVPD SAQTSNTHTT SASPPEETEL QAM

35
GENBANK ID: P17774
VERSION P17774 GI:121324

40
1 MSESIVVCDV AEDLVEKLRK FRFRKETNNA AIIMKIDKDK RLVLDEELE GISPDELKDE
61 LPERQPRFIV YSYKYQHDDG RVSYPCLFIF SSPVGCKPEQ QMMYAGSKNK LVQTAELTKV
121 FEIRNTEDLT EEWLREKLG FH

45
GENBANK ID: P51858
VERSION P51858 GI:1708157

50
1 MSRSNRQKEY KCGDLVFAKM KGYPHWPARI DEMPEAAVKS TANKYQVFFF GTHETAFLGP
61 KDLFPYEESK EKFGKPNKRK GFSEGLWEIE NNPTVKASGY QSSQKKSCVE EPEPEPEAAE
121 GDGDKKGNAE GSSDEEGKLV IDEPAKEKNE KGALKRRAGD LLEDSPKRPK EAENPEGEEK
181 EAATLEVERP LPMEVEKNST PSEPGSGRGP PQEEEEEEDE EEEATKEDAE APGIRDHESL
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55
GENBANK ID: P10147
VERSION P10147 GI:127078

60
1 MQVSTAALAV LLCTMALCNQ FSASLAADTP TACCFSYTSR QIPQNFADY FETSSQCSKP
61 GVIFLTRSR QVCADPSEEW VQKYVDLEL SA

65
GENBANK ID: P13500
VERSION P13500 GI:126842

1 MKVSAALLCL LLIAATFIPQ GLAQPDAINA PVTCCYNFTN RKISVQRLAS YRRITSSKCP
61 KEAVIFKTIV AKEICADPKQ KVVQDSMDHL DKQTQTPKT

GENBANK ID: NP_065391.1
VERSION NP_065391.1 GI:10092621

1 MGVLLTQRTL LSLVLALLFP SMASMAAIGS CSKEYRVLLG QLQKQTDLMQ DTSRLLDPI

5
61 RIQGLDVPKL REHCRERPGA FPSEETLRGL GRRGFLQTLN ATLGCVLHRL ADLEQRLPKA
121 QDLERSGLNI EDLEKLQMAR PNILGLRNNI YCMAQLLDNS DTAEPTKAGR GASQPPTPTP
181 ASDAFQRKLE GCRFLHGYHR FMHSVGRVFS KWGESPNRSR RHSPHQALRK GVRRTSPSRK
241 GKRLMTRGQL PR

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GENBANK ID: XP_013053.3
VERSION XP_013053.3 GI:14768277

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1 MEKERETLQA WKERVQQLD RVVAFWMEHS HDQEHGGFFT CLGREGRVYD DLKYVWLQGR
61 QVWMYCRLYR TFERFRHAQL LDAAKAGGEF LLRYARVAPP GKKCAFVLTR DGRPVKVQRT
121 IFSECFYTM MNELWRATGE VRYQTEAVEM MDQIVHWVQE DASGLGRPQL QGAPAAEPMA
181 VPMMLNLVE QLGEADEELA GKYAELGWC ARRILQHVQR DGQAVLENVS EGGKELPGCL
241 GRQONPGHTL EAGWFLLRHC IRKGDPFLRA HVIDKFLLLP FHSGWDPDHG GLFYFQDADN
301 FCPTQLEWAM KLWWPHSEAM IAFLMGYSDS GDPVLLRLFY QVAEYTFRQF RDPEYGEWFG
361 YLSREGKVAL SIKGGPFKGC FHVPRCLAMC EEMLGALLSR PAPAPSPAPT PACRGAE

20
GENBANK ID: B31848
VERSION B31848 GI:87005

25
1 MTCKMSQLER NIETIINTFH QYSVKLGHPD TLNQGEFKEL VRKDLQNFLK KENKNEKVIE
61 HIMEDLDTNA DKQLSFEEFI MLMARLTWAS HEKMHEGDEG PGHHHKPGLG EGTP

30
GENBANK ID: CAA38698.1
VERSION CAA38698.1 GI:35522

35
1 MPVMRLFPCF LQLLAGLALP AVPPQOWALS AGNGSSEVEV VPFQEVWGRS YCRALERLVD.
61 VVSEYPSEVE HMFSPSCVSL LRCTGCCGDE NLHCVPVETA NVTMQLLKIR SGDRPSYVEL
121 TFSQHVRCEC RPLREKMKPE RCGDAVPRR

40
GENBANK ID: AAA35789.1
VERSION AAA35789.1 GI:181971

45
1 MNFLLSWVHW SLALLLYLHH AKWSQAAPMA EGGGQNHHEV VKFMDVYQRS YCHPIETLVD
61 IFQEYPDEIE YIFKPSCVPL MRCGGCCNDE GLECVPTES NITMQIMRIK PHQGOHIGEM
121 SFLQHNKCEC RPKKDRARQE NPCGPCSERR KHLFVQDPQT CKCSCKNTDS RCKARQLELN
181 ERTCRCDKPR R

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GENBANK ID: AAA66062.1
VERSION AAA66062.1 GI:536898

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1 MWKRWLALAL ALVAVAVVRA EEELRSKSKI CANVFCGAGR ECAVTEKGEP TCLCIEQCKP
61 HKRPVCGSNG KTYLNHCELH RDACLTGSKI QVDYDGHCKE KKSVPSPASP VVCYQSNRDE
121 LRRRIQWLE AEIIPDGWFS KGSNYSEILD KYFKNFDNGD SRLDSSEFLK FVEQNETAIN
181 ITTYPDQENN KLLRGLCVDA LIELSDENAD WKLSFOEFLK CLNPSFNPPE KKCALEDETY
241 ADGAETEVD C NRCVCACGNW VCTAMTCDGK NQKGAQTQTE EEMTRYVQEL QKHQETAECT
301 KRVSTKEI

60
GENBANK ID: AAA59872.1
VERSION AAA59872.1 GI:398038

65
1 MGWLPLLLLLL TQCLGVPGQR SPLNDFQVLR GTELQHLLHA VVPGPWQEDV ADAEECAGRC
61 GPLMDCRAFH YNVSSHGCQL LPWTOHSPHT RLRRSGRCDL FQKKDYVRTC IMNNGVGYRG
121 TMATTVGGLP CQAWSHKFPN DHKYTPTLRN GLEENFCRNP DGDPPGPGWY TTDPAVRFQS
181 CGIKSCREAA CVWCNGEYR GAVDRTEGSR ECQRWDLQHP HQHPFEPGKF LDQGLDDNYC
241 RNPDGSERPW CYTTDPQIER EFCDLPRCGS EAQPRQEATT VSCFRGKGEY YRGTAANTTA
301 GVPCQRWDAQ IPHQHRTPE KYACKDLREN FCRNPDGSEA PWCFTLRPGM RAAFCYQIRR
361 CTDDVRPQDC YHGAGEQYRG TVSKTRKGVO CORWSAETPH KPQFTFTSEP HAQLEENFCR
421 NPDGDSHGPW CYTMDPRTPE DYCALRRCAD DQPPSILDPP DQVQFEKCGK RVDRLDQRRS
481 KLRVVGHPG NSPWTVSLRN RQGOHFCGGS LVKEQWILTA RQCFSSCHMP LTGYEVWLGT
541 LFQNPQHGEF SLQRVFVAKM VCGPSGSQLV LLKLEERSVTL NORVALICLP PEWYVVPFGT
601 KCEIAGWGET KGTGNDTVLN VAFNLVISNQ ECNIKHRGRV RESEMCTEGL LAPVGACEGD
661 YGGPLACFTH NCWVLEGIII PNRVCARSRW PAVFTRVSF VDWIHKVMRL G

GENBANK ID: P01579
VERSION P01579 GI:124479

1 MKYTSYILAF QLCIVLGS LG CYCQDPYVKE AENLK KYFNA GHSDVADNGT LFLGILKNWK
61 EESDRKIMQS QIVSFYFKLF KNFKDDQSIQ KSVETIKEDM NVKFFNSNKK KRDDFEKLTN
121 YSVTDLNVQR KAIHELIQVM AELSPA AKTG KRKRSQMLFR GRRASQ

5 GENBANK ID: XP_035842.1
DEFINITION HOMO SAPIENS SMALL INDUCIBLE CYTOKINE A5 (RANTES) (SCYA5), MRNA.

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121 TGCTTTGCCT ACATTGCCCG CCCACTGCCC CGTGCCACAC TCAAGGAGTA TTTCTACACC
181 AGTGGCAAGT GCTCCAACCC AGCAGTCGTC TTTGTCAACC GAAAGAACCG CCAAGTGTGT
15 241 GCCAACCAG AGAAGAAATG GGTTCGGGAG TACATCAACT CTTTGGAGAT GAGCTAGGAT
301 GGAGAGTCCT TGAACCTGAA CTTACACAAA TTTGCCTGTT TCTGCTTGCT CTTGTCCTAG
361 CTTGGGAGGC TTCCCTCAC TATCCTACCC CACCCGCTCC TTGAAGGGCC CAGATTCTAC
421 CACACAGCAG CAGTTACAAA AACCTTCCCC AGGCTGGACG TGGTGGCTCA CGCCTGTAAT
481 CCCAGCACTT TGGGAGGCCA AGGTGGGTGG ATCACTTGAG GTCAGGAGTT CGAGACCAGC
20 541 CTGGCCAACA TGATGAAACC CCATCTCTAC TAAAAATACA AAAAATTAGC CGGGCGTGGT
601 AGCGGGCGCC TGTAATCCCA GCTACTCGGG AGGCTGAGGC AGGAGAAATGG CGTGAACCCG
661 GGAGGCGGAG CTTGCAGTGA GCGGAGATCG CGCCACTGCA CTCCAGCCTG GGCGACAGAG
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781 GGCCACGCC TGTAATCCCA GCTACTCGGG AGGCTAAGGC AGGAAAATTG TTTGAACCCA
25 841 GGAGGTGGAG GCTGCAGTGA GCTGAGATTG TGCCACTTCA CTCCAGCCTG GGTGACAAAG
901 TGAGACTCCG TCACAACAAC AACCAACAAA AGCTTCCCCA ACTAAAGCCT AGAAGAGCTT
961 CTGAGGCGCT GCTTTGTCAA AAGGAAGTCT CTAGGTTCTG AGCTCTGGCT TTGCCTTGGC
1021 TTTGCCAGGG CTCTGTGACC AGGAAGGAAG TCAGCATGCC TCTAGAGGCA AGGAGGGGAG
1081 GAACACTGCA CTCTTAAGCT TCCGCCGTCT CAACCCCTCA CAGGAGCTTA CTGGCAAACA
30 1141 TGAAAATCG G

GENBANK ID: CAA72079.1
DEFINITION H.SAPIENS MRNA FOR ESTROGEN SULFOTRANSFERASE.

CDS 63..947
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61 CAATGAATTC TGAAGTGGAC TATTATGAAA AGTTTGAAGA AGTCCATGGG ATTCTAATGT
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40 181 TTGTCATTGC CACCTACCCT AAATCTGGTA CAACCTGGGT TAGTGAAATT GTGTATATGA
241 TCTATAAAGA GGGTGATGTG GAAAAGTGCA AAGAAGATGT AATTTTTAAT CGAATACCTT
301 TCCTGGAATG CAGAAAAGAA AACCTCATGA ATGGAGTAAA ACAATTAGAT GAGATGAATT
361 CTCCTAGAAT TGTGAAGACT CATTGCCCAC CTGAACTTCT TCCTGCCTCA TTTTGGGAAA
421 AGGATTGTAA GATAATCTAT CTTTGCCGGA ATGCAAAGGA TGTGGCTGTT TCCTTTTATT
45 481 ATTTCTTCT AATGGTGGCT GGTATCCAA ATCCTGGATC CTTTCCAGAG TTTGTGGAGA
541 AATTCATGCA AGGACAGGTT CCTTATGGTT CCTGGTATAA ACATGTAAA TCTTGGTGGG
601 AAAAGGGAAA GAGTCCACGT GTACTATTTT TTTTCTACGA AGACCTGAAA GAGGATATCA
661 GAAAAGAGGT GATAAAATTG ATACATTTCC TGGAAAGGAA GCCATCAGAG GAGCTGTGG
721 ACAGGATTAT ACATCACTT TCGTTCCAAG AGATGAAGAA CAATCCATCC ACAAATTACA
50 781 CAACACTGCC AGACGAAATT ATGAACCAGA AATTGTCGCC CTTTATGAGA AAGGGAATTA
841 CAGGAGACTG GAAAAATCAC TTTACAGTAG CCCTGAATGA AAAATTTGAT AAACATTATG
901 AGCAGCAAAT GAAGGAATCT AACTGAAGT TTCGAAGTGA GATCTAAGAA GGTCTT

GENBANK ID: AAA63210.1
DEFINITION HUMAN KERATINOCYTE GROWTH FACTOR MRNA, COMPLETE CDS.

CDS 446..1030
/CODON_START=1

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61 TCTGCTGGAG AACTTTTCAG CTGAGAAATA GTTTGTAGCT ACAGTAGAAA GGCTCAAGTT
121 GCACCAGGCA GACAACAGAC ATGGAATTCT TATATATCCA GCTGTTAGCA ACAAACAAA
181 AGTCAAATAG CAAACAGCGT CACAGCAACT GAACTTACTA CGAACTGTTT TTATGAGGAT
241 TTATCAACAG AGTTATTTAA GGAGGAATCC TGTGTTGTTA TCAGGAACTA AAAGGATAAG
301 GCTAACAATT TGGAAAGAGC AAGTACTCTT TCTTAAATCA ATCTACAATT CACAGATAGG
65 361 AAGAGGTCAA TGACCTAGGA GTAACAATCA ACTCAAGATT CATTTTCATT ATGTTATTCA

5 421 TGAACACCCG GAGCACTACA CTATAATGCA CAAATGGATA CTGACATGGA TCCTGCCAAC
481 TTTGCTCTAC AGATCATGCT TTCACATTAT CTGTCTAGTG GGTACTATAT CTTTAGCTTG
541 CAATGACATG ACTCCAGAGC AAATGGCTAC AAATGTGAAC TGTTCCAGCC CTGAGCGACA
601 CACAAGAAAGT TATGATTACA TGGGAAGGAGG GGATATAAGA GTGAGAAGAC TCTTCTGTCTG
661 AACACAGTGG TACCTGAGGA TCGATAAAAG AGGCAAAGTA AAAGGGACCC AAGAGATGAA
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781 GGTGGAAAGT GAATTCTATC TTGCAATGAA CAAGGAAGGA AAACCTCTATG CAAAGAAAGA
841 ATGCAATGAA GATTGTAAC TCAAAGAACT AATTCTGGAA AACCATTACA ACACATATGC
10 901 ATCAGCTAAA TGGACACACA ACGGAGGGGA AATGTTTGT GCCTTAAATC AAAAGGGGAT
961 TCCTGTAAGA GGAAAAAAA CGAAGAAAGA ACAAAAACA GCCCACTTTC TTCTATGGC
1021 AATAACTTAA TTGCATATGG TATATAAGA ACCCAGTTCC AGCAGGGAGA TTTCTTTAAG
1081 TGGACTGTTT TCTTCTTCT CAAAATTTTC TTTCTTTTA TTTTCTAGTA ATCAAGAAAG
1141 GCTGGAAAAA CTAAGTGAAG ACTGATCAAG CTGGACTTGT GCATTTATGT TTGTTTAAAG
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15 1261 TTGTAAAAAA TTGTAAACT GGTGTACAA TCATGATGTT AGTAACAGTA ATTTTCTCT
1321 TAAATTAATT TACCCTTAAG AGTATGTTAG ATTTGATTAT CTGATAATGA TTATTTAAAT
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1441 GGTATATCAG ACCTACAGGC TTCTGGCAGG ATTTGTCAGA TAATCAAGCC AACTAACTA
1501 TGGAAATGA GCAGCATTTT AAATGCTTTC TAGTGAAAAA TTATAATCTA CTTAACTCT
20 1561 AATCAGAAAA AAAATCTCA AAAAACTAT TATGAAAGTC AATAAAATAG ATAATTTAAC
1621 AAAAGTACAG GATTAGAACA TGCTTATACC TATAAATAAG AACAAAATTT CTAATGCTGC
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1741 CAGTGATTGA TGATAATACT GTACTTCATC TTACTTGCCA CAAAATAACA TTTTATAAAT
25 1801 CCTCAAAGTA AAATTGAGAA ATCTTTAAGT TTTTTCAGG TAACATAATC TATCTTTGTA
1861 TAATTCATAT TTGGGAATAT GGCTTTTAAAT AATGTTCTTC CCACAAATAA TCATGCTTTT
1921 TTCCTATGGT TACAGCATTA AACTCTATTT TAAGTTGTTT TTGAACTTTA TTGTTTGTG
1981 ATTTAAGTTT ATGTTATTTA TAAAAAAA ACCTTAATAA GCTGTATCTG TTTTATATGC
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30 2101 TGAAGTACAC TAAGTCTAGC ACACAGCACT TGGGCCAGCA AATCCTGGAA GCAGACAAAA
2161 ATAAGAGCCT GAAGCAATGC TTACAATAGA TGTCTCACAC AGAACAATAC AAATATGTAA
2221 AAATCTTTTC ACCACATATT CTTGCCAATT AATTGGATCA TATAAGTAAA ATCATTACAA
2281 ATATAAGTAT TTACAGGATT TTAAGTTAG AATATATTTG AATGCATGGG TAGAAAATAT
2341 CATATTTTAA AACTATGTAT ATTTAAATTT AGTAATTTTC TAATCTCTAG AAATCTCTGC
35 2401 TGTTCAAAAG GTGGCAGCAC TGAAAGTTGT TTTCTGTTA GATGGCAAGA GCACAATGCC
2461 CAAAATAGAA GATGCAGTTA AGAATAGGG GCCCTGAATG TCATGAAGGC TTGAGGTCAG
2521 CCTACAGATA ACAGGATTAT TACAAGGATG AATTCTCACT TCAAAAGTCT TTCATTGGCA
2581 GATCTTGGTA GCACTTTTATA TGTTTACCAC TGGGAGGTCA ATATTTATCT AATTTAAAG
2641 GTATGCTAAC CACTGTGGTT TTAATTTCAA AATATTTGTC ATTCAAGTCC CTTTACATAA
40 2701 ATAGTATTTG GTAATACATT TATAGATGAG AGTTATATGA AAAGGCTAGG TCAACAAAAA
2761 CAATAGATTC ATTTAATTTT CCTGTGGTTG ACCTATACGA CCAGGATGTA GAAACTAGA
2821 AAGAACTGCC CTTCTCAGA TATACTCTTG GGAGAGAGCA TGAATGGTAT TCTGAAGTAT
2881 CACCTGATTC AAGGACTTTG CTAGCTAGGT TTTGAGGTCA GGCTTCAGTA ACTGTAGTCT
2941 TGTGAGCATA TTGAGGGCAG AGGAGGACTT AGTTTTTCAT ATGTGTTTCC TTAGTGCTTA
45 3001 GCAGACTATC TGTTTATAAT CAGTTTTCAG TGTGAATTCA CTGAATGTTT ATAGACAAAA
3061 GAAAATACAC ACTAAACTA ATCTTCATTT TAAAAGGGTA AAACATGACT ATACAGAAAT
3121 TTAAATAGAA ATAGTGTATA TACATATAAA ATACAAGCTA TGTTAGGACC AAATGCTCTT
3181 TGTCTATGGA GTTATACTTC CATCAAATTA CATAGCAATG CTGAATTAGG CAAAACCAAC
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50 3301 GACTTTTAAAT TATTTGTTTT TCTCCTATTT TTAAATTTAT TATGCAAATT TTAGAAAATA
3361 AAATTTGCTC TAGTTACACA CCTTTAGAA TCTAGAATAT TAAAAGTGA AGGGGCCTCC
3421 ATCCCTCTTA CTCATTTGTA GTCTAGGAAA TTGAGATTTT GATACACCTA AGGTCACGCA
3481 GCTGGGTAGA TATACAGCTG TCACAAGAGT CTAGATCAGT TAGCACATGC TTTCTACTCT
3541 TCGATTATTA GTATTATTAG CTAATGGTCT TTGGCATGTT TTTGTTTTTT ATTTCTGTG
55 3601 AGATATAGCC TTTACATTTG TACACAAATG TGAATATGTC TTGGCAATGC ACTTCATACA
3661 CAATGACTAA TCTATACTGT GATGATTTGA CTCAAAAGGA GAAAAGAAAT TATGTAGTTT
3721 TCAATTCTGA TTCCTATTCA CTTTTGTTT ATGAATGGAA AGCTTTGTGC AAAATATACA
3781 TATAAGCAGA GTAAGCCTTT TAAAATGTT CTTTGAAAGA TAAATTTAAA TACATGAGTT
3841 TCTAACAATT AGA

60 GENBANK ID: AAA62202.1
DEFINITION HUMAN ENDOTHELIAL-MONOCYTE ACTIVATING POLYPEPTIDE II MRNA, COMPLETE
CDS.
VERSION U10117.1 GI:498909
MRNA 1..1057
35 CDS 50..988
/CODON_START=1

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121 TCTTAAGCAG CAAGTTTCTC TACTTAAGGA GAAAGCAATT TTGCAGGCAA CTTTGAGGGA
181 AGAGAAGAAA CTTCGAGTTG AAAATGCTAA ACTGAAGAAA GAAATTGAAG AACTGAAACA
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961 AACCATGAGC AACAGTGGAA TCAAATAAAA TGCTTCCACT ACCAAAAGAC ATTAGAGAAA
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GENBANK ID: P17936

DEFINITION HUMAN ACIDIC FIBROBLAST GROWTH FACTOR MRNA, 5' END, CLONE
LAMBDA-MJ36.

CDS 358..>478
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121 AGGACAAGTG GATCCAACAG CCTTCGCTCC AGGGGAATCA GGGCATCGCC TCCTTTTCTG
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241 CAGCTGTCCT GGTAGAACAG TGTGGACATT GCAGAAGCTG TCACTGCCCC AGAAAGAAAG
301 CACCCACAGAG CCAAGGCAA GAGTCTTGAA AGCGCCACAA GCAGCAGCTG CTGAGCCATG
361 GCTGAAGGGG AAATCACCAC CTTACAGGCC CTGACCGAGA AGTTTAATCT GCCTCCAGGG
421 AATTACAAGA AGCCCAAACCT CCTCTACTGT AGCAACGGGG GCCACTTCCT GAGGATCC

GENBANK ID:

U76376.1

VERSION U76376.1 GI:1923234

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GENBANK ID: Y00638

VERSION Y00638.1 GI:34280

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30
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HHSTSYNSKALIAFLAFLIIVTSIALLVLYKIYDLHKKRSCNLDEQQELVERDDEKQ
LMNVEPIHADILLETYKRKIADEGRF LAEFQSI PRVFSKFPIKEARKPFNQKNRYV
DILPYDYNRVELSEINGDAGSNYINASYIDGFEKEPRKYIAAQGPRDETVDDEFWRMIWE
QKATVIVMVTRCEEGRNRKCAEYWFSMEEGTRAFGDVVVKINQHKRCPDYIIQKLNIV
NKKEKATGREVTHIQFTSWPDHGVFPEDPHLLLKLRRRVNAFNSFFSGPIVVHCSAGVG
RTGTYIGIDAMLEGLEAENKVDVYGVVVKLRRQRCLMVQVEAQYILIHQALVEYNQFG
ETEVNLSELHPYLHNMKKRDPPSESPLEAEFQRLPSYRSWRTQHIGNQEENKSKNRN
SNVIPYDYNRVPLKHELEMSKEHSDSDSDSDSEEPSKYINASFIMSYWKPEVM
IAAQGPLKETIGDFWQMI FQRKVIVMLTELKHGDQEICAQYWGEKGQTYGDIQVLDL
KDTDKSSTYTLRVFELRHSKRKDSRTVYQYQYTNWSVEQLPAEPKELISMIQVVKQKL

PQKNSSEGNKHHKSTPLLIHCRDGSQQTGIFCALLNLLSAETEEVVDIFQVVKALRK
ARLGMVSTFEQYQFLYDVIASSTYPAQNGQVKNNHQEDKIEFDNEVDKVKQDANCNP
LGAPEKLPEAKEQAEGSEPTSGTEGPEHSVNGPASPALNQGS

5 GENBANK ID: AF001383.1
VERSION AF001383.1 GI:2199534

10 MAEMGSKGVTAGKIASNVQKKLTRAQEKVLQKLGADETKDEQF
EQCVQNFNKLTEGTRLQKDLRTYLASVKAMHEASKKLNELQEVYEPDWPGRDEANK
IAENNDLLWMDYHQKLVLDQALLTMDTYLGQFPDIKSRIAKRGRKLVYDSARHHYESL
QTAKKKDEAKIAKEEELIKAQKVFEEMNVDLQEELPSLWNSRVGFYVNTFQSIAGLE
ENFHKEMSKLNQNLNDVLVGLEKQHGNTFTVKAQPSDNAPAKGNKSPSPDGSPAAT
PEIRVNHEPEPAGGATPGATLPKSPSQLRKGPVPPPKHTPSKEVKQEQLSLFEDT
15 FVPEISVTTSPQAEASEVAGGTQPAAGAQEPGETAASEAASSSLPAVVVETFPATVN
GTVEGGSGAGRLDLPFGFMFKVQAQHDYTATDTDELQKAGDVVLVIPPQNPEEQDEG
WLMGVKESDWNQHELEKCRGVFPENFTERVP

20 GENBANK ID: XM_038595.3
VERSION XM_038595.3 GI:18590923

25 MAPPSEETPLIPQRSCSLLSTEAGALHVLLPARGPGPPQRLSFS
FGDHLAEDLCVQAAKASGILPVYHSLFALATEDLSCWFPPSHIFSVEDASTQVLLYRI
RFYFPNWFGLEKCHRFGRLKDLASAILDLPVLEHLFAQHRSDLVSGRLPVGLSLKEQG
ECLSLAVLDLARMAREQAORPGELLKTVSYKACLPPSLRDLIQGLSFVTRRRIRRTVR
RALRRVAACQADRHSIMAKYIMDLERLDPAGAAETFHVGLPGALGGHDGLGLLRVAGD
GGIAWTQGEQEVLPFCDFPEIIVDISIKQAPRVGPAGEHRLVTVTRTDNQILEAEFPG
LPEALSFVALVDGYFRLTTDSQHFFCKEVAPRLLLEEVAEQCHGPITLDFAINKLKTG
GSRPGSYVLRSPQDFDSFLTVCVQNPLGPDYKGLIRRSPTGTFLLVGLSRPHSSL
30 RELLATCWDGGLHVDGVAVTLTSCCIPRPKEKSNLIIVQRGHSPPTSSLVQPSQYQL
SQMTFHKIPADSLEWHENLHGGSFTKIYRGRHEVVDGEARKTEVLLKVMDAKHKNCM
ESFLEAASLMSQVSRYHLVLLHGVCMAGDSTMVQEFVHLGAIDMYLRKRGLVPASWK
LQVVKQLAYALNYLEDKGLPHGNVSARKVLLAREGADGSPFIKLSDPGVSPAVLSLE
MLTDRIPWVAPECLREAQTLSEADKWGFATVWEVFSGVTMPISALDPAKKLQFYED
35 RQQLPAKWTETALLIQQCMAYEPVQRPFRVIRDLNSLISSDYELLSDPGALAP
RDGLWNGAQLYACQDPTIFEERHLKYISQLGKGNFGSVELCRYDPLGNTGALVAVKQ
LQHSQPDQQRDFQREIQILKALHSDFIVKYRGVSYGPGRQSLRLVMEYLPSCGLRDFL
QRHRRLDASRLLYSSQICKGMEYLGSRRCVHRDLAARNILVESEAHVKIADFGGLAK
LLPLDKDYVVREPGQSPIFWYAPESLSDNIFSRQSDVWSFGVVLVYELFTYCDKSCSP
40 SAEFLRMMGCERDVPALCRLELEEGQRLPAPPACPAEVHELMKLCWAPSPQDRPSF
SALGPQLDMLWSGSRGCETHAFTAHEGKHHSLSFS

45 GENBANK ID: M32292.1
VERSION M32292.1 GI:181492

50 MENSLRCVWVPKLAFLVFGASLLSAHLQVTGFQIKAFATLRFLS
EPSDAVTMRGGNVLLDCAESDRGVFVIKWKDGIHLALGMDERKQQLSNGSLLIQNI
LHSRHHKPDEGLYQCEASLGDGSIISRTAKVAVAGPLRFLSQTESVTAFMGDTVLLK
CEVIGEPMPPTIHWQKNQDQLTPIPGDSRVVLPAGALQISRLQPGDIGIYRCSARNPA
SSRTGNEAEVRILSDPGLHRQLYFLQRPNSVVAIEGKDAVLECCVSGYPPPSFTWLRG
EEVIQLRSKKYSLGGSNLLISNVTDDSGMYTCVVITYKNENISASAELTVLVPPWFL
NHPSNLYAYESMDIEFECTVSGKPVPTVNMKNGDVVIPSDFQIVGGSNLRILGVVK
SDEGFYQCVAENEAGNAQTSACLIVPKPAIPSSSVLPSAPRDVVPVLVSSRFVRLSWR
PPAEAKGNIQTFTVFFSREGDNRRERLNTTQPGSLQLTVGNLKPAMYTRVVAAYNEW
35 GPGESSQPIKVATQPELQVPGPVENLQAVSTSPTSILITWEPPAYANGPVQGYRLFCT
EVSTGKEQNIQVGLSYKLEGLKFTESLRLFLAYNRYGPGVSTDDITVVTLSDVPSA
PPQNVSLVNSRSIKVSWLPPPSGTQNGFITGYKIRHRKTTRGEMETLEPNNLWYL
FTGLEKGSQYSFQVSAMTVNGTGPSPNWTAEPTENDLDESQVDPQPSLHVRPQTNC
IIMSWTPPLN

60 GENBANK ID: X06318
VERSION X06318.1 GI:35488

65 MADPAAGPPPSEGEESTVRFARKGALRQKNVHEVKNHKFTARFF
KQPTFCSHCTDFIWGFGKQGFQCCFVVKRCHEFVTFCPGADKGPASDDPRSKH
KFKIHTYSSPTFCDHCGSLLYGLIHQGMKCDTCMMNVHKRCVMNVPSLCGTDHTEERRG

5 RIYIOAHIDRDVLIVLVRDAKNLVPMDPNGLSDPYVKLKLIPDPKSESKQKTKTIKCS
LNPEWNETFRFQLKESDKDRRLSVEIWDWDLTSRNDFMGSLSGISELQKASVDGWFK
LLSQEEGEYFNVPPVPEGSEANEELRQKFERAKISQGTKVPEEKTNTVSKFDNNGNR
DRMKLTDFNFLMVLGKGSFGKVMLSERKGTDELYAVKILKKDVVIQDDDVECTMVEKR
VLALPGKPPFLTQLHSCFQTMDRLYFVMEYVNGGDLMYHIQQVGRFKEPHAVFYAAEI
AIGLFFLQSKGIIYRDLKLDNVMLDSEGHKIADFGMCKENIWDGVTTKTFCGTPDYI
APEIIAYQPYGKSVDWWAFGVLLYEMLAGQAPFEGEDEDELQFSIMEHNVAYPKSMK
EAVAICKGLMTKHPGKRLGCGPEGERDIKEHAFFRYIDWEKLERKEIQPPYKPKARDK
10 RDTSNFDKEFTRQPVLETPDKLFIMNLDQNEFAGFSYTNPEFVINV

GENBANK ID: J04132.1
VERSION J04132.1 GI:623041

15 MKWKALFTAAILQAQLPITEAQSFGLLDPKLCYLLDGILFIYGV
ILTALFLRVKFSRSAEPPAYQOQONQLYNELNLGRREEYDVLDRRGRDPFEMGGKPRR
KNPQEGLYNELQKDKMAEAYSEIGMKGERRRGKGHDGLYQGLSTATKDTYDALHMQAL
PPR

GENBANK ID: U04313.1
VERSION U04313.1 GI:453368

20 MDALQLANSFAVDLQKQCEKEPLGNVLFSPICLSTSLSLAQV
GAKGDTANEIGQVLHFENVKDIFFGFQTVTSDVNKLSSFYSLKLIKRLYVDKSLNLST
25 EFISSTKRPYAKELETVDKDKLEETKGQINNSIKDLTDGHFENILADNSVNDQTKIL
VVNAAYFVGKWMKKFPESETKECPFRLNKTDTPVQMMNEATFCMGNIDSINCKIIE
LPFQNKHLSMFILLPKDVEDESTGLEKIEKQLNSESLSQWTNPSTMANAKVKLSIPKE
KVEKMIDPKACLENLGLKHIFSEDTSDFSGMSETKGVALSNIHKVCLEITEDGGDSI
EVPGARILQHKDELNADHPFIYIIRHNKTRNIIFFGKFCSP

30 GENBANK ID: X68968.1
VERSION X68968.1 GI:452315

35 MDEPSPLAQPLELNQHSRFFIGSVSEDNSEDEISNLVKLDLLEE
KEGSLSPASVGSDDLSDGISSLDGLALHIRSSMSGHLHVKQGRDRKKIDSQRDFTV
ASPAEFVTRFGGNKVIEKVLIANNGIAAVKCMRSIRRWSYEMFRNERAIRFVVMVTPE
DLKANAAYIKMADHYVPVPGGPNNNYANVELILDIAKRIPVQAVWAGWGHASENPKL
PELLLNKNGIAFMGPPSQAMWALGDKIASSIVAQTAGIPTLPWSGSLRVDWQENDFSK
RILNVPQELYEKGYVKDVEDGLQAAEEVGYPVMIKASEGGGGKGIRKVNADDFFPNLF
RQVQAEVPGSPIFVMRLAKQSRHLEVQILADQYGNALISLEGRDCSVQRRHQKIEEAP
40 ATIATPAVFEHMEQCAVKLAKMVGYSAGTVEYLYSQDRSFYFLELNPRLQVEHPCTE
MVADVNLPAAQLQIAMGIPLYRIKDIRMYGVSPWGDSPIDFEDSAHVPCPRGHVIAA
RITSENPDGFKPSSGTQVELNFRSNKNVWGYFSVAAAGGLHEFADSQFGHCFWGES
REEAISNMVVALKELSIRGDFRTTVEYLKLETESFQMNRI DTGWLDRLIAEKVQAE
RPDTMLGVVCGALHVADVSLRNSVSNFLHSLERGQVLPALHTLNTVDVELIYEGVKYV
45 LKVTRQSPNSYVIMNGSCVEVDVHRLSDGGLLSYDGSSYTTMKEEVDYRITIGN
KTCVFEKENDPSVMRSPSAGKLIQYIVEDGGHVLAGQCYAEIEVMKMVMTLTAVESGC
IHYVKRPGAALDPGCVLAKMQLDNPSKVQQAELHTGSLPRIQSTALRGEKLRHVFHYV
LDNLVNMNGYCLPDPFSSSKVKDWVERLMKTLRDPSPLELQDIMTSVSGRIPPV
EKSICKEMAQYASNITSVLQCFPSQQIANILDSHAATLNRKSEREVFFMNTQSIVQLV
50 QRYRSGIRGHMKAVVMDLLRQYL RVETQFQNGHYDKCVFALREENKSDMNTVLNIF
HAQVTKKNLLVTMLIDQLCGRDPTLTDELLNLTTELTLQSKTTNAKVALRARQVLIA
HLPSYELRHNQVESIFLSAIDMYGHQFCIENLQKLILSETSI FDVLPNFFYHSNQVVR
MAALEVYVRRAYIAYELNSVQHRQLKDNTCVVEFQFMLPTSHPNRGNIPTLNRMSFSS
NLNHYGMTHVASVSDVLLDNSFTPPCQRMGMVSRFTFEDFVRI FDEVMGCFSDSPPQ
55 SPTFPEAGHTSLYDEDKVPRDEPIHILNVAIKTDCDIEDDLAAMFREFTQONKATLV
DHGIRRLTFLVAQKDFRKQVNYEVDRRFHREFPKFFTFRARDKFEEDRIYRHLEPALA
FQLELNRMRNFDLTAIPCANHKMHLYLGAKEVGTETDYRFFVRAIRHSDLVTK
ASFEYLQNEGERLLEAMDELEVAFNNTNVRTDCNHIFLNFVPTVIMDPSKIEESVRY
MVMRYGSRWLKRLVLAQEVKINIRQTTGSAVPIRLFITNESGYLDISLYKEVTD
60 SGNIMFHSFGNKQGPQHGLINTPYVTKDLLQAKRFQAQTLGTTYIYDFPEMFRQALF
KLWGSPPDKYPKDILTYTELVLDSQGLVEMNRLPGGNEVGMVAFKMRFKTQEYPEG
VIVIGNDITFRIGSFGPGEDLLYLRASEMARAIAIPKIYVAANS GARIGMAEEIKHMF
HVAWVDPEDPHKGFKYLYLTPQDYTRISSLSNVHCKHIEEGGESRYMITDIIGKDDGL
GVENLRGSGMIAGESSLAYEEIVTISLVTFCRAIGIGAYLVRGQRVIQVENSIIITG
65 ASALNKVLGREVYTSNNQLGGVQIMHYNGVSHITVPDDFEGVYTILEWLSYMPKDNHS
VPPIITPTDPIDREIEFLPSRAPYDPRWMLAGRPHTLKG TWQSGFFDHGSFKEIMAP

5 WAQTVVTGRARLGGIPVGVIIVETRTVEVAVPADPANLDSEAKIIQQAGQVWFPSAY
KTAQAIKDFNREKLPLMIFANWRGFSGGMKDMDQVLKFGAYIVDGLRQYKQPIIYI
RPMRELRGSSWVIDATINPLCIEMYADKESRGGVLEPEGTVEIKFRKEDLIKSMRRI
DPAYKKLMEQLGEPDLSDKDRKDLLEGRLKAREDLLLPYHQVAVQFADFHDTPGRMLE
KGVISDILEWKTARTFLYWRRLRLLEDQVKQEILQASGELSHVHIQSMRLRRWFVETE
GAVKAYLWDNNQVQVWLEQHWQAGDGPRSTIRENITYLKHDSVLKTIRGLVEENPEV
AVDCVIYLSQHISPAERAQVHVHLLSTMDSPAST

10 GENBANK ID: L03840.1
VERSION L03840.1 GI:182570

15 MRLLLALLGVLLSVPGPPVLSLEASEEVELEPCLAPSLEQQEQE
LTVALGQPVRLCCGRAERGWHYKEGSRLAPAGRVRGWRGRLEIASFLPEDAGRYLCL
ARGSMIVLQNLTLITGDSLTSSNDDDEPKSHRDPNSRHSYPQAPYWTHPORMEKKLH
AVPAGNTVKFRCPAAGNPTPTIRWLKDGQAFHGENRIGGIRLRHQHWSLVMSVPSD
RGTYTCLVENAVGSIRYNYLLDVLESPHRPILQAGLPANTTAVVGS DVELLCKVYS
AQPHIQWLKHIVINGSFGADGFYVQVLTADINSSEVEVLYLRNVSAEDAGEYTCL
AGNSIGLSYQSAWLTVPEDPTWTAAPEARYTDIILYASGSLALAVLLLAGLYRG
QALHGRHPRPPATVQKLSRFPLARQFSLESSESGKSSSSLVRGVRLSSSGPALLAGLV
20 SLDLPLDPLWEFPRDRVLGKPLGEGCFGQVVRAEAFGMDPARPDQASTVAVKMLKDN
ASDKDLADLVSEMEVMKLGHRKNIINLLGVCTQEGPLYVIVECAAKGNLREFLRARR
PPGPDLS PDGPRSSSEGPLSFVLVSCAYQVARGMOYLESRKCIHRDLAARNVLVTEDN
VMKIDAFGLARGVHHIDYKKTSGRLPVKWMPEALFDRVYTHQSDVWSFGILLWEI
FTLGGSPYPGIPVEELFSLREGHRMDRPPHCPPELYGLMRECWAAPSQRPTFKQLV
25 EALDKVLLAVSEEYDLRLTFGPYSPSGGDASSTCSSSDSVFSDPLPLGSSSFPPGS
GVQT

30 GENBANK ID: AF043342.1
VERSION AF043342.1 GI:2905633

VRSSSRTPSDKPVAVVANPQAEQQLQWLNRRANALLANGVELR
DNQLVVPSEGLYLIYSQVLFKGQGCPSHVLTLHTISRIAVSYQTKVNLLSAIKSPCQ
RETFRGAEPWYEPYILGGVFQLEKGDRLSAEINRPDYLDFAESGQVYFGIIAL

35 GENBANK ID: NM_000735.2
VERSION NM_000735.2 GI:10800407

MDYYRKYAAIFLVTLSVFLHVLHSA PDVQDCPECTLQENPFFSQ
40 PGAPILQCMGCCFSRAYPTPLRSKKTMLVQKNVTSESTCCVAKSYNRVTVMGGFKVEN
HTACHCSTCYHKS

45 GENBANK ID: M83533.1
VERSION M83533.1 GI:178541

LRKHNIETYLIKQPEDSLLSLPEDIVKESVSSSDRRNSGATFTE
GSWSPELPFDNIVGKQNTLAALTRNSINLLPNHLAQALHVQSGPEEINKRIEHTIDLR
SGDKLRREHIKPFSLMFKDSSLEHKYSQMRDEVFKSNLVCAFIVLLFITAIQSLPSS
RVMPMTIQFSILIMLHSAVLITTAEDYKCLPLILRKTCCWINETYLARNVIFASIL
INFLGAILNIIWCFDKSIPLKNLTFNSSAVFTDICSYPEYFVFTGVLMVTCVFLR
50 LNSVLKLAVLLIMIAIYALLTETVYAGLFLRYDNLNHSGEDFLGTKEVSLLLMMAMFL
AVFYHGQQLLEYTARLDFLRVQAKEEINEMKELREHNENMLRNILPSHVARHFLEKDR
DNEELYSQSYDAVGVMFASIPGFADFYSQTEMNNOGVECLRLLEIIADFDELLGEDR
FQDIEKIKTIGSTYMAVSGLSPEKQCEDKWGHLALADFSALATESIQEINKHSFNN
FELRIGISHGSSVAGVIGAKKPOYDIWGKTVNLASRMDSTGVSGRIQVPEETYLILKD
55 QGFADFYRGEIYVKGISEQEGKIKTYFLLGRVQPNPFILPPRRLPGQYSLAAVVLGLV
QSLNRQROKQLNENNTGIIKGHYNRRTLSPSGTEPGAQEGTDKSDLP

60 GENBANK ID: M31767.1
VERSION M31767.1 GI:181615

MDKDCEMKRTTLDSPGLKLELSGCEQGLHEIKLLGKGTSAADAV
EVPAPAAVLGGPEPLMQCTAWLNAYFHQPEAIEEFVPALHHPVFQOESFTRQVLWKL
LKVVKEGEVISYQQLAALAGNPKATRAVGGAMRGNPVPILPCHRVVCSSGAVGNYS
GLAVKEWLLAHEGHRGKPLGGSSGLAGAWLKGAGATSGSPAGRN

65 GENBANK ID: X14723

VERSION X14723.1 GI:30250

5 MMKTLLLFVGLLLTWESGQVLGDQTVSDNELQEMSNQGSKYVNK
EIQNAVNGVKQIKTLIEKTNEERKTLLSNLEEAKKKKEDALNETRESETKLKELPGVC
NETMMALWEECKPCLKQTCMKFYARVCRSGSLVGRQLEEFNLQSSPFYFWMNGDRID
SLENDRQQTHMLDVMQDHFSRASSIIDELFQDRFTREPQDTYHYLPFSLPHRRPHF
FFPKSRIVRSIMPFSPYEPNLFHAMFQPFLEMIHEAQQAMDIHFHSPAFOHPPTFIR
EGDDDRTVCREIRHNSTGCLRMKDQCDKREILSVDCSTNNPSQAKLRRELDDESLOVA
10 ERLTRKYNELLKSYQWKMLNTSSILLEQLNEQFNWVSRLANLTQGEDQYYLRVTTVASH
TSDSDVPSGVTEVVVKLFSDPITVTVPEVSRKNPKFMETVAEKALQEYRKKHREE

GENBANK ID: X04391.1
VERSION X04391.1 GI:37186

15 MPMGSLQPLATLYLLGMLVASCLGRLSWYDPDFQARLTRSNSKC
QGQLEVYLKDGWHMVCSQSWGRSSKQWEDPSQASKVCQRLNCGVPLSLGPFVLYTTPQ
SSIICYGQLGSFSNCSHSRNDMCHSLGLTCLPQKTPPTTRPPPTTTPEPTAPPRLO
LVAQSGGQHCAGVVEFYSGSLGGTISYEAQDKTQDLENFLCINNLCGSFLKHLPETEA
GRAQDPGEPREHQPLPIQWKIQNSSCTSLHCFRKKIPQKSGRVLALLCSGFQPKVQS
20 RLVGSSICEGTVEVRQGAQWALCDSSSARSSLRWEEVCREQQCGSVNSYRVLDAGD
PTSRGLFCPHQKLSQCHELWERNYSYCKKVFVTCQDPNPAGLAAGTVASIIILALVLLV
LLVVCPLAYKKLVKKFRQKQWIGPTGMNQNMFSHRNHTATVRSHAENPTASHVD
NEYSQPPRNSRLSAYPALEGVLHRSSMQPDNSSDSYDLHGAQRL

25 GENBANK ID: S78187.1
VERSION S78187.1 GI:243485

30 MEVPQPEPAPGSALSPAGVCGGAQRFGHLPGLLLGSHGLLGSPV
RAAASSPVTTLTQTMHDLAGLSRSRLTHLSLSRRASESSLSSESSESSDAGLCMDSP
SPMDPHMAEQTFEQAIQAASRIIRNEQFAIRRFQSMFVRLGHSPVLRNITNSQAPDG
RRKSEAGSGAASSSGEDKENDGFVKMPWKPTHSPSTHALAEWASRREAFQRPSSAP
DLMCLSPDRKMEVEELSPLALGRFSLTPAEGDTEEDDGFVDILESDLKDDDAVPPGME
SLISAPLVKTLKEKEEKDLVMYSKQRLFRSPMPCSVIRPILKRLERPQDRDTPVQN
KRRRSVTPPEEQEAEKPKARLSKSLCHDEIENLLDSHRELIGDYSKAFLLQTV
35 GKHQDLKYISPETMVALLTGKFSNIVDKFVIVDCRYPYEYEGGHIKTAVNLPLERDAE
SFLKSPIAPCSLDKRVILIFHCFESSERGPRMCRFIRERDRAVNDYPSLYYPEMYIL
KGGYKEFFPQHPNFCPEQDYRPMNHEAFKDELKTRFKTRSWAGERSRRELCDRLODQ

40 GENBANK ID: Y00096.1
VERSION Y00096.1 GI:30455

45 MREAAMYSTAVAI FLVILVAALQGSAPRESPLPYHIPLDPEG
LELSWNVSYTQEAIFQLLVRLKAGVLFMSDRGELENADLVVLWTDGDTAYFADAW
SDQKGQIHLDPQDYQLLQVQRTPEGLTLLFKRPFQTCDPKDYLIEDGTVHLVYGILE
EPFRSLEAINSGSLQMLQVQLLKNIPPELPSDTCTMEVQAPNIQIPSQETTYWC
YIKELPKGFSRHHIIKYEPIVTKGNEALVHHMEVFQCAPEMDSVPHFSGPCDSKMKPD
RLNYCRHVLAAWALGAKAFYYPEAGLAFGGPGSSRYLRLEVHYHNPLVIEGRNDSSG
IRLYYTAKLRRFNAGIMELGLVYTPVMAIPPRETAFILTGYCTDKCTQLALPPSGIHI
FASQLHHTLTGRKVTVLVRDGREWEIVNQDNHYSHPHFEIRMLKKVSVHFGDVLIT
50 SCTYNTEDRELATVGGFGILEEMCVNYVHYYPQTQLELCKTAVDAGFLQKYFHLINRF
NNEDVCTCPQASVSQQTSPVWNSFNCVLLKALYSFAPISMHCNKSSAVRFQGEWNLQ
PLPKVISTLEEPTQCPTSQGRSPAGPTVVSIGGGKG

55 GENBANK ID: XM_055551.3
VERSION XM_055551.3 GI:18557356

60 MKETQKSTYYITGESKEQVANSFAVERVRKQGFEVVMTEPIDE
YCVQQLKEFDGKSLVSVTKGLELPEDEEEKKKMEESKEKFENLCKLMKEILDKKVEK
VTISNRLVSSPCCIVTSTYGTANMEQIMKAQALRDNSTMGYMAKKHLEINPDHPIM
ETLRQKAEADKNDKAVKDLVLLFETALLSSGFSLEDQTHSNHIYHMIKGLGLTDED
EVAAEPPSDAVPDEIPPLEGDEDEDASRMEEVD

65 GENBANK ID: M84711.1
VERSION M84711.1 GI:182774

5 MAVGKNKRLTKGGKKGAKKKVVDPFSSKKDWYDVKAPAMFNIRNI
GKTLVTRTQGTKIASDGLKGRVFEVSLADLQNDVAFRKFKLITEDVQGNCLTNFHG
MDLTRDKMCSMVKKWQTMIEAHVDVKTDDGYLLRLFCVGFTHKRNQIRKTSYAQHQQ
VRQIRKKMEIMTREVQTNDLKEVVNKLIPDSIGKDIEKACQSIYPLHDVFVRKVKML
KKPKFELGKLMELHGESSSGKATGDETGAVERADGYEPPVQESV

10 GENBANK ID: X53505
VERSION X53505.1 GI:36145

MAEEGIAAGGVMDVNTALQEVLTALIHDLARGIREAAKALDK
RQAHLCVQASNCDEPMYVKLVEALLAEHQINLIKVDNKKLGEWVGLCKIDREGNPRK
VVGCSQVVKDYGKESQAKDVIEEYFKCKK

15 GENBANK ID: X06617
VERSION X06617.1 GI:36143

MADIQTERAYQKOPTIFQNKRRVLLGETGKEKLPRYYKNIGLGF
KTPKEAIEGTYIDKKCPFTGNVSIRGRILSGVVTKMKMORTIVIRRDYLHYIRKYNRF
EKRHKNSVHLSPCFRDVQIGDIVTVGECRPLSKTVRFNLKVTKAAGTKKQFQKF

20 GENBANK ID: M55040.1
VERSION M55040.1 GI:177974

MRPPQCLLHTPSLASPLLLLLLWLLGGGVGAEGREDAELLVTVR
25 GGRLRGIRLKTGGPVSAFLGIPFAEPPMGPRRFLPPEPKQPSGVVDATTFQSVCYQ
YVDTLYPGFEGTEMWNPNNRELSDECLYNVWTPYPRPTSPTPVLVWIYGGGFYSGASS
LDVYDGRFLVQAERTVLVSMNYRVGAFGLALPGSREAPGNVGLLDQRLALQWQENV
AAFGGDPSTVTLFGESAGAASVGMHLLSPSRGLFHRAVLQSGAPNGPWATVGMGEAR
RRATQLAHLVGCPPGGTGGNDTELVACLRTRPAQVLVNHEWHVLPQESVFRFSFVPV
30 DGDFLSDTPEALINAGDFHGLQVLVGVVKDEGSYFLVYGAPGFSKDNESLISRAEFLA
GVRVGVVPQVSDLAEEAVVLHYTDWLHPEDPARLREALSDVVDHNVVCPVAQLAGRLA
AQGARVYAYVFEHRASTLSWPLWMGVPHGYEIEFIFGIPLDPSRNYTAEKI FAQRLM
RYWANFARTGDPNEPRDPKAPQWPPYTAGAQYVSLDLRPLEVRRGLRAQACAFWNRF
LPKLLSATDTLDEAERQWKAEFHRWSSYMHVWKNQFDHYSKQDRCSDL

35 GENBANK ID: NM_000717.2
VERSION NM_000717.2 GI:9951925

MRMLLALLALSAAEPSASAESHWCYEVQAESSNYPCLVPVKWGG
40 NCQKDRQSPINIVTTKAKVDKKLGRFFFSGYDKKQTTWTQVNNGHSMMLLENKASISG
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DEIAVLAFLVEAGTQVNEGFOPLVEALSNIKPEMSTTMAESSLLDLLPKEEKLRHYF
RYLGSLTTPTEDEKVVTVFREPIQLHREQILAFSQKLYDKEQTVSMKDNVRPLQQL
45 QORTVIKSGAPGRPLPWALPALLGPMACLLAGFLR

GENBANK ID: S70587.1

VERSION S70587.1 GI:546848

50 MTALFLMSMLFGLACGQAMSFCIPTETMHIERRECAVCLTINT
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55 GENBANK ID: M34057
VERSION M34057.1 GI:339547

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60 LCQIPVHGASVPKLYQHSQQPGKALGTHVIHSTHTLPLTVTSQQGVKVKFPPNIVNIH
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PHVYPVAAKTQLGRCFQETIGSQCGKALPGLSKQEDCCGTVGTSWGFNKCQKCPKKPS
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PGTAAFKEICPGMGYTVSGVHRRRPIHHHVKGKPVFVKPKNTQPVAKSTHPPPLPAK
65 EEPVEALTFSREHGARSAPPEVATAPPEKEIPSLDQEKTKLEPGQPQLSPGISAIHLH
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5 VRYTCICYEGYRFSEQQRKCVDI DECTOVQHLCSQGRCENTEGSFLCICPAGFMASEE
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10 HRHLCAHGQCRNTEGSFQCVCDQGYRASGLGDHCEINEDKSVQCQRGDCINTAGS
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CEDIDECVNNTVCDSHGFCNTAGSFRCLCYQGFQAPQDGGQCVDVNECELLSGVCGE
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RDALVDFSEQYTPADPYFIQDRFLNSFEELQAECEGILNGCENGRCVRVQEGYTCDC
15 LDGYHLDTAKMTCFDVNECDELNNRMSLCKNAKCINTDGSYKCLCLPGYVPSDKPNYC
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20 GENBANK ID: AF257099.1
VERSION AF257099.1 GI:8037944

MSDAAVDTSSSEITTEDLKEKKEVVEEAENGRDAPAHGNANEENG
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EDD

25 GENBANK ID: L06505.1
VERSION L06505.1 GI:186799

MPPKFDPNKIKVYLRCTGGEVGATSALAPKIGPLGLSPKKVGD
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30 ECPAS

35 GENBANK ID: X79234.1
VERSION X79234.1 GI:495125

MAQDQGEKENPMRELRIKLCNICVGESGGRLTRAQVLEQLT
GQTPVFSKARYTVRSFGIRRNEKIAVHCAVRGAKAEIILEKGLKVRELELRKNNFSDT
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40 GENBANK ID: X59932.1
VERSION X59932.1 GI:30255

MSAIQAAWPSGTECIAKYNFHGTAEQDLFPCKGDVLTIVAVTKD
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45 LFLVRESTNYPGDYTLVCSDGKVEHYRIMYHASKLSIDEEVYFENLMQLVEHYTSDA
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50 KDVVPRVEKGYKMDAPDGCPPAVYEVMMKNCWHLDAAMRPSFLQLREQLEHIKTHELHL

55 GENBANK ID: AAA98616.1
VERSION AAA98616.1 GI:178428

1 MQGPWVLLLL GLRLQLSLGI IPVEEENPDF WNRQAAEALG AAKKLOPAQT AAKNLIMFLG
61 DGMGVSTVTA ARILKGQKKD KLGPEFTLAM DRFPYVALSK TYSVDKHVPD SGATATAYLC
60 121 GVKGNFQTIG LSAAARFNQC NTTRGNEVIS VVNRKAKAGK SVGVVTTTRV QHASPAGTYA
181 HTVNRNWYSD ADVPASARQE GCQDIATQLI SNMDIDVILG GGRKYMFPMPG TPDPEYPDDY
241 SQGGTRLDGK NLVQEWLAKH QGARYVWNRT ELLQASLDPS VTHLMGLFEP GDMKYEIHRD
301 STLDPSIMEM TEAALLLSR NPRGFFLEVE GGRIDHGHHE SRAYRALTET IMFDDAIERA
361 GQLTSEEDTL SLVTADHSHV FSFGGYPLRG SSIFGLAPGK ARDRKAYTVL LYGNPGGYVL
65 421 KDGARPDVTE SESGSPEYRQ QSAVPLDGET HAGEDVAVFA RGPQAHLVHG VQEQTFAHV
481 MAFAACLEPY TACDLAPPAG TTDAAHGPS VVPALLPLA GTLLLLGTAT AP

5 GENBANK ID: XM_041507.1
VERSION XM_041507.1 GI:14737457

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10 VTAIIFCVALSAYDLVLAEDEEMNRMHESMKLFDSICNNKWFTDTSIILFLNKKDLFE
EKITHSPLTICFPEYTGANKYDEAASYIQSKFEDLNKRKDTKEIYTHFTCATDTKNVQ
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15 GENBANK ID: NM_001032.2
VERSION NM_001032.2 GI:13904868

MGHQQLYWSHPRKFGQGSRSRVCNHRHGLIRKYGLNMCRCFR
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20 GENBANK ID: M22430.1
VERSION M22430.1 GI:190888

MKTLLLLAVIMIFGLLQAHGNLVNFHRMIKLTGKEAALS YGFY
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25 CRSQCECDKAAATCFARNKTTYNKYQYYSNKHCRGSTPRC

GENBANK ID: X63527
VERSION X63527.1 GI:36127

30 MSMLRLQKRLASSVLRCGKKKVWLDPNETNEIANANSRQQIRKL
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RRLRLRYRESKKIDRHYHSLYLVKGNVFNKRILMEHIHKLKADKARKKLLADQAE
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35 GENBANK ID: AF099644.1
VERSION AF099644.1 GI:4323527

MAQFAFESDLHSLQLDAPINAPPARWQRKAKEAAGPAPSPMR
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40 NSQTPKKEHQKAWALNLNGFDVEEAKILRLSGKPQNAPEGYQNRKLVLSQKATPGS
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NSYILSSGSRSGHIIHHHDVRAEHHVATLSGHSQEVCGLRWAPDGRHLASGNDNLVN
VWPSAPGEGGWVPLQTFTHQGAVKAWCPWQSNVLTGGGTSDRHIRIWNVCSGAC
45 LSAVDAHSQVCSILWSPHYKELISGHGFAQNQLVIWKYPTMAKVAELKGHTSRVLSLT
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GENBANK ID: X51466
VERSION X51466.1 GI:31105

50 MVNFTVDQIRAIMDKKANIRNMSVIAHVDHGKSTLTDSLVCAG
IIASARAGETRFTDTRKDEQERCITIKSTAISLFYELSENDLNFQSKDGAGFLINL
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55 ATSPEGKKLPRTFCQLILDPIFKVFDAIMNFKKEETAKLIEKLDIKLDESDKDKEGKP
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PLMMYISKMVPTS DKGRFYAFGRVFSGLVSTGLKVRIMGPNYTPGKKEDLYLKPIQRT
ILMMGRYVEPIEDVPCGNIVGLVGVDQFLVKTGTITTFEHAHNMRVMKFSVSPVVRVA
VEAKNPADLPKLVEGLKRLAKSDPMVQCIIEESGEHIIAGAGELHLEICLDLEEDHA
60 CIPIKKSDPVVSYRET VSESNVCLSKSPNKHNRLYMKARFPDGLAEDIDKGEVSA
RQELKQARYLAEKYEWDAEARKIWCFGPDGTGNILTDITKGVQYLNEIKDSVVAG
FQWATKEGALCEENMRGVRFVDVHDLHADAIHRGGGQIIPARRCLYASVLTAQPRL
MEPIYLVETIQCEQVVGGIYGLNRKRGHVFEESSQVAGTFMFVVKAYLPVNESFGFTA
65 DLRSTGGQAFPCVFDHWQILPGDPFDNSSRPSQVVAETRKRLKGLKEGIPALDNFLD

GENBANK ID: M15661
VERSION M15661.1 GI:337577

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MVNVPKTRRTFCKKCGKHQPHKVTQYKKGKDSLYAQGRRRYDRK
QSGYGGQTKPIFRKKAATTKIVLRLECVEPNCRSKRMLAIKRCKHFELGGDKKRKGQ
VIQF

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GENBANK ID: J04823.1
VERSION J04823.1 GI:1311703

MSVLTPLLLRGLTGSARRLPVPRAKIHSLPPEGKLGIMELAVGL
TSCFVTFLPAGWILSHLETYRRPE

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GENBANK ID: NM_001760.2
VERSION NM_001760.2 GI:16950657

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MELLCCETRHRAPRAGPDPRLLGDQRLVLSLLRLEERYVPRASY
FQCVQREIKPHMRKMLAYWMLEVCEEQRCCEEVFLAMNYLDLYLSCVPTRKAQLQLL
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GPSQTSTPTDVTAIHL

25

GENBANK ID: NM_002625.1
DEFINITION HOMO SAPIENS 6-PHOSPHOFRUCTO-2-KINASE/FRUCTOSE-2, 6-BIPHOSPHATASE 1
(PFKFB1), MRNA.
VERSION NM_002625.1 GI:4505744
CDS 80..1495

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1 GAATTCCGGA CAGGTAGTAA GATAGGAAGT GAGGCCAGGT ACCTTGTGGG CAGTGATGTC
61 ATTCGGTGCG ACTCCTAAGA TGTCTCCAGA GATGGGAGAG CTCACCCAAA CCAGGTTGCA
121 GAAGATCTGG ATTCCACACA GCAGCGGCAG CAGCAGGCTG CAACGGAGAA GGGGCTCATC
181 CATACCCAG TTTACCAATT CCCCCACAAT GGTGATCATG GTGGGTTTAC CAGCTCGAGG
241 CAAGACCTAT ATCTCCACAA AGCTCACACG ATATCTCAAC TGGATAGGAA CACCAACTAA
301 AGTGTTTAAT TTAGGCCAGT ATCGACGAGA GGCAGTGAGC TACAAGAACT ATGAATTCTT
361 TCTTCCAGAC AACATGGAAG CCCTGCAAAT CAGGAAGCAG TCGCCCTGG CAGCCCTGAA
421 GGATGTTTAC AACTATCTCA GCCATGAGGA AGGTCATGTT GCGGTTTTTG ATGCCACCAA
481 CACTACCAGA GAACGACGGT CACTGATCCT GCAGTTTGCA AAAGAACATG GTTACAAGGT
541 GTTTTTTATT GAGTCCATTT GTAATGACCC TGGCATAATT GCAGAAAACA TCAGGCAAGT
601 GAAACTTGGC AGCCCTGATT ATATAGACTG TGACCGGGAA AAGGTTCTGG AAGACTTTCT
661 AAAGAGAATT GAGTGCTATG AGGTCAACTA CCAACCCCTG GATGAGGAAC TGGACAGCCA
721 CCTGTCTTAC ATCAAGATCT TCGACGTGGG CACACGCTAC ATGGTGAACC GAGTGCAGGA
781 TCACATCCAG AGCCGCACAG TCTACTACCT CATGAATATC CATGTACACAC CTCGCTCCAT
841 CTACCTTTGC CGACATGGCG AGAGTGAAC CAACATCAGA GGCCGCATCG GAGGTGACTC
901 TGGCCTCTCA GTTCGCGGCA AGCAGTATGC CTATGCCCTG GCCAACTTCA TTCAGTCCCA
961 GGGCATCAGC TCCCTGAAGG TGTGGACCAG TCGCATGAAG AGGACCATCC AGACAGCTGA
1021 GGCCCTGGGT TCCCTTATG AGCAGTGGAA GGCCCTGAAT GAGATTGATG CGGGTGTCTG
1081 TGAGGAGATG ACCTATGAAG AAATCCAGGA ACATTACCCT GAAGAATTTG CACTGCGAGA
1141 CCAAGATAAA TATCGCTACC GCTATCCCAA GGGAGAGTCC TATGAGGATC TGGTTCAGCG
1201 TCTGGAGCCA GTGATAATGG AGCTAGAACG ACAGGAGAAT GTACTGGTGA TCTGCCACCA
1261 GGCTGTCATG CGGTGCCTCC TGGCCTATTT CCTGGATAAA AGTTCAGATG AGCTTCCATA
1321 TCTCAAGTGC CCTCTGCACA CAGTGCTCAA ACTCACTCCT GTGGCTTATG GCTGCAAAGT
1381 GGAATCCATC TACCTGAATG TGGAGGCGGT GAACACACAC CGGGAGAAGC CTGAGAATGT
1441 GGACATCACC CGGGAACCTG AGGAAGCCCT GGATACTGTC CCAGCCCACT ACTGAGCCCT
1501 TTCCAAGAAG TCAAACCTGC TGTGTCTTCA TCGCCTTCCA CCTTTAGGAA ATGCTATCTT
1561 TGCTCTTCTC CTACTCTGCC TTGGCCTCAC TGAGGCACCC CACTTCCAGT GAAGAAGTCC
1621 TCCGCAACTC CCAAACAAGC CTCGCTTGCT GGCCGCAACC AAGGAGCTAT CTAGCTCTGG
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1741 G

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GENBANK ID: D00760
DEFINITION HUMAN MRNA FOR PROTEASOME SUBUNIT HC3.
VERSION D00760.1 GI:220023
CDS 1..705
/CODON_START=1

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1 ATGGCGGAGC GCGGGTACAG CTTTTGCTG ACTACATTCA GCCCGTCTGG TAAACTTGTC
61 CAGATTGAAT ATGCTTTGGC TGCTGTAGCT GGAGGAGCCC CGTCCGTGGG AATTAAAGCT
121 GCAAATGGTG TGGTATTAGC AACTGAGAAA AAACAGAAAT CCATTCTGTA TGATGAGCGA
181 AGTGATACACA AAGTAGAACC AATTACCAAG CATATAGGTT TGGTGTACAG TGGCATGGGC
241 CCCGATTACA GAGTGCTTGT GCACAGAGCT CGAAAACCTAG CTCAACAATA CTATCTTGTC
301 TACCAAGAAC CCATTCCTAC AGCTCAGCTG GTACAGAGAG TAGCTTCTGT GATGCAAGAA
361 TATACTCAGT CAGGTGGTGT TCGTCCATTT GGAGTTTCTT TACTTATTTG TGGTTGGAAT
421 GAGGGACGAC CATATTTATT TCAGTCAGAT CCATCTGGAG CTTACTTTGC CTGGAAAGCT
481 ACAGCAATGG GAAAGAACTA TGTGAATGGG AAGACTTTCC TTGAGAAAAG ATATAATGAA
541 GATCTGGAAC TTGAAGATGC CATTACATACA GCCATCTTAA CCCTAAAGGA AAGCTTTGAA
601 GGGCAAATGA CAGAGGATAA CATAGAAGTT GGAATCTGCA ATGAAGCTGG ATTTAGGAGG
661 CTTACTCCAA CTGAAGTTAA GGATTACTTG GCTGCCATAG CATAACAATG AAGTGACTGA
721 AAAATCCAGA ATTTTCAGATA ATCTATCTAC TTAAACATGT TTAAAGTATG TTTTGTGTTG
781 CAGACTTTTT GCATACTTAT TTCTACATGG TTAAATCGA CTGTTTTTAA AATGACACTT
841 ATAAATCCTA ATAAACTGTT AAACCC

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GENBANK ID: P10644

GENBANK ID: XM_043948.2

DEFINITION HOMO SAPIENS ALDOLASE A, FRUCTOSE-BISPHOSPHATE (ALDOA), MRNA.

VERSION XM_043948.2 GI:18585537

CDS 243..1349

/CODON_START=1

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40
45
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121 ACCCCTTTCC TTCCCACAGG TCCCTGGCCA AAGATTTATT TCTCTTGACA ACCAAGGGCC
181 TCCGTCTGGA TTTCCAAGGA AGAATTTCTT CTGAAGCACC GGAAGTGGCT ACTACCAGCA
241 CCATGCCCTA CCAATATCCA GCACTGACCC CGGAGCAGAA GAAGGAGCTG TCTGACATCG
301 CTCACCGCAT CGTGGCACCT GGCAAGGGCA TCCTGGCTGC AGATGAGTCC ACTGGGAGCA
361 TTGCCAAGCG GCTGCAGTCC ATTGGCACCG AGAACACCGA GGAGAACCGG CGCTTCTACC
421 GCCAGCTGCT CTGACAGCT GACGACCCG TGAACCCCTG CATTGGGGGT GTCATCCTCT
481 TCCATGAGAC ACTCTACCAG AAGGCGGATG ATGGGCGTCC CTTCCCCCAA GTTATCAAAT
541 CCAAGGGCGG TGTGTGGGC ATCAAGGTAG ACAAGGGCGT GGTCCCCCTG GCAGGGACAA
601 ATGGCGAGAC TACCACCCAA GGGTTGGATG GGCTGTCTGA GCGCTGTGCC CAGTACAAGA
661 AGGACGGAGC TGAATTCGCC AAGTGGCGTT GTGTGCTGAA GATTGGGGAA CACACCCCTT
721 CAGCCCTCGC CATCATGGAA AATGCCAATG TTCTGGCCCG TTATGCCAGT ATCTGCCAGC
781 AGGTGGGCTT GCAGAATGGC ATTGTGCCCA TCGTGGAGCC TGAGATCCTC CCTGATGGGG
841 ACCATGACTT GAAGCGCTGC CAGTATGTGA CCGAGAAGGT GCTGGCTGCT GTCTACAAGG
901 CTCTGAGTGA CCACCACATC TACCTGGAAG GCACCTTGCT GAAGCCCAAC ATGGTCACCC
961 CAGGCCATGC TTGCACTCAG AAGTTTTCTC ATGAGGAGAT TGCCATGGCG ACCGTCACAG
1021 CGCTGCGCCG CACAGTGCCC CCCGCTGTCA CTGGGATCAC CTTCTGTCTT GGAGGCCAGA
1081 GTGAGGAGGA GCGGTCCATC AACCTCAATG CCATTAACAA GTGCCCCCTG CTGAAGCCCT
1141 GGGCCCTGAC CTTCTCCTAC GGCCGAGCCC TGCAAGCCCTC TGCCCTGAAG GCCTGGGGCG
1201 GGAAGAAGGA GAACCTGAAG GCTGCGCAGG AGGAGTATGT CAAGCGAGCC CTGGCCAACA
1261 GCCTTGCCCTG TCAAGGAAAG TACACTCCGA CGGGTCAGGC TGGGGCTGCT GCCAGCGAGT
1321 CCCTCTTCGT CTCTAACCAC GCCTATTAAG CGGAGGTGTT CCCAGGCTGC CCCAACACT
1381 CCAGGCCCTG CCCCCTCCCA CTCTTGAAGA GGAGGCCGCC TCCTCGGGGC TCCAGGCTGG
1441 CTTGCCCGCG CTCTTTCTTC CCTCGTGACA GTGGTGTGTG GTGTCGTCTG TGAATGCTAA
1501 GTCCATCACC CTTTCCGGCA CACTGCCAAA TAAACAGCTA TTTAAGGGGG

55
GENBANK ID: NM_005175.1

DEFINITION HOMO SAPIENS ATP SYNTHASE, H+ TRANSPORTING, MITOCHONDRIAL FO COMPLEX, SUBUNIT C (SUBUNIT 9), ISOFORM 1 (ATP5G1), MRNA.

VERSION NM_005175.1 GI:4885080

CDS 120..530

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121 TGCAGACCGC CGGGGCATTA TTCATTTCTC CAGCTCTGAT CCGCTGTTGT ACCAGGGGTC
181 TAATCAGGCC TGTGTCTGCC TCCTTCTTGA ATAGCCAGT GAATTCATCT AAACAGCCTT
241 CCTACAGCAA CTTCCCACTC CAGGTGGCCA GACGGGAGTT CCAGACCAGT GTTGTCTCCC
301 GGGACATTGA CACAGCAGCC AAGTTTATTG GTGCTGGGGC AGCCACAGTT GGTGTGGCTG

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361 GTTCAGGGGC TGGCATTGGA ACCGTGTTTG GCAGCTTGAT CATTGGCTAT GCCAGGAACC
421 CGTCTCTCAA GCAGCAGCTC TTCTCCTATG CCATTCTTGG CTTTGCCCTG TCTGAGGCCA
481 TGGGGCTTTT CTGTTTGATG GTCGCCTTCC TCATCCTCTT CGCCATGTGA GGCTCCATGG
541 GGGGTCACCG GCCTGTTGCT ACTGCAACTC CACACCATTG TTGGTGCTGG GGTGTGTTAA
601 GCTTTACCAT TAAACACAAC GTTCTCTTAA A

GENBANK ID: M20496.1

DNA LINEAR

DEFINITION HUMAN CATHEPSIN L GENE, COMPLETE CDS.

VERSION M20496.1 GI:809235

CDS 134..1135

/CODON_START=1

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1 ACCTCCACGT GCCCTGTTTT TCTGGAGGCA CATCCTTGGC CTCTTCCACA GTCCTTGGGT
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181 AGCTACTCTA ACATTTGATC ACAGTTTAGA GGCACAGTGG ACCAAGTGGG AGGCGATGCA
241 CAACAGATTA TACGGCATGA ATGAAGAAGG ATGGAGGAGA GCAGTGTGGG AGAAGAACAT
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361 CATGAACGCC TTTGGAGACA TGACCAAGTGA AGAATTCAGG CAGGTGATGA ATGGCTTCA
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481 ATCTGTGGAT TGGAGAGAGA AAGGCTACGT GACTCCTGTG AAGAATCAGG GTCAGTGTGG
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661 AGGCTGCAAT GGTGGCCTAA TGGATTATGC TTTCCAGTAT GTTCAGGATA ATGGAGGCCT
721 GGACTCTGAG GAATCCTATC CATATGAGGC AACAGAAGAA TCCTGTAAGT ACAATCCCAA
781 GTATTCTGTT GCTAATGACA CCGGCTTTGT GGACATCCCT AAGCAGGAGA AGGCCCTGAT
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901 CCTGTTCTAT AAAGAAGGCA TTTATTTTGA GCCAGACTGT AGCAGTGAAG ACATGGATCA
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961 TGGTGTGCTG GTGTTGGCT ACGGATTTGA AAGCACAGAA TCAGATAACA ATAAATATTG
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1261 AATTCTGTGA TATTTTCACA CTGGTAAATG TTACCTCTAT TTTAATTACT GCTATAAATA
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1381 TTTTACCTG TTTAAATAAA ATCG

GENBANK ID: XM_031596.3

DEFINITION HOMO SAPIENS ANNEXIN A4 (ANXA4), MRNA.

VERSION XM_031596.3 GI:18553329

CDS 48..770

/CODON_START=1

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1 GAAGAACTTC TGCTTGGGTG GCTGAACTCT GATCTTGACC TAGAGTCATG GCCATGGCAA
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181 ACCGCAACAC CGCCAGCGC CAGGAGATCA GGACAGCCTA CAAGAGCACC ATCGGCAGGG
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361 GCACTGATGA GGGCTGCCTA ATTGAGATCC TGGCCTCCCG GACCCCTGAG GAGATCCGGC
421 GCATAAGCCA AACCTACCAG CAGCAATATG GACGGAGCCT TGAAGATGAC ATTGCTCTG
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841 CGATGATAAC ACCCTCATCA GAGTGATGGT TTCTCGAGCA GAAATTGACA TGTGATAT
901 CCGGGCACAC TTCAAGAGAC TCTATGAAA GTCTCTGTAC TCGTTCATCA AGGGTGACAC
961 ATCTGGAGAC TACAGGAAAG TACTGCTTGT TCTCTGTGGA GGAGATGATT AAAATAAAAA
1021 TCCCAGAAGG ACAGGAGGAT TCTCAACACT TTGAATTTT TTAACCTCAT TTTTCTACAC
1081 TGCTATTATC ATTATCTCAG AATGCTTATT TCCAATTAAA ACGCCTACAG CTGCCTCCTA
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1141 GAATATAGAC TGTCTGTATT ATTATTCACC TATAATTAGT CATTATGATG CTTTAAAGCT
1201 GTACTTGCAT TTCAAAGCTT ATAAGATATA AATGGAGATT TTAAAGTAGA AATAAATATG

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1261 TATTCCATGT TTTTAAAAGA TTACTTTCTA CTTTGTGTTT CACAGACATT GAATATATTA
1321 AATTATTCCA TATTTTCTTT TCAGTGAAAA ATTTTAAATA TGGAAAGACTG TTCTAAAATC
1381 ACTTTTTTCC CTAATCCAAT TTTTAGAGTG GCTAGTAGTT TCTTCATTG AAATTGTAAG
1441 CATCCGGTCA GTAAGAATGC CCATCCAGTT TTCTATATTT CATAGTCAAA GCCTTGAAAG
1501 CATCTACAAA TCTCTTTTTT TAGGTTTTGT CCATAGCATC AGTTGATCCT TACTAAGTTT
1561 TTCATGGGAG ACTTCCTTCA TCACATCTTA TGTGAAATC ACTTTCTGTA GTCAAAGTAT
1621 ACCAAAACCA ATTTATCTGA ACTAAATCTT AAAGTATGGT TATACAAACC ATATACATCT
1681 GGTACCAAAA CATAAATGCT GAACATTCCA TATTATTATA GTTAATGTCT TAATCCAGCT
1741 TGCAAGTGAA TGGAAAAAAA AATAAGCTTC AAAGTAGGTA TTCTGGGAAT GATGTAATGC
1801 TCTGAATTTA GTATGATATA AAGAAAACCT TTTTGTGCTA AAAATACTTT TTAATAATCA
1861 TTTTGTGAT GTAGTAATT TCTATTGCA CTGTGCCTTT CAACTCCAGA AACATTCTGA
1921 AGATGTACTT GGATTTAATT AAAAAGTTCA CTTTGT

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GENBANK ID: M22865.1
DEFINITION HUMAN CYTOCHROME B5 MRNA, COMPLETE CDS.
VERSION M22865.1 GI:181226

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53..457
CODON_START=1
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1 CAGCCAGCTC GACGGGGCTG TGTGTGCTGG GCCTGGCTCG CGGCGAACCG AGATGGCAGA
61 GCAGTCGGAC GAGGCCGTGA AGTACTACAC CCTAGAGGAG ATTCAGAAGC ACAACCACAG
121 CAAGAGCACC TGGCTGATCC TGCACCACAA GGTGTACGAT TTGACCAAAT TTCTGGAAGA
181 GCATCCTGGT GGGGAAGAAG TTTTAAAGGA ACAAGCTGGA GGTGACGCTA CTGAGAACTT
241 TGAGGATGTC GGGCACTCTA CAGATGCCAG GGAAATGTCC AAAACATTCA TCATTGGGGA
301 GCTCCATCCA GATGACAGAC CAAAGTTAAA CAAGCCTCCG GAAACTCTTA TCACTACTAT
361 TGATTCTAGT TCCAGTTGGT GGACCAACTG GGTGATCCCT GCCATCTCTG CAGTGGCCGT
421 CGCCTTGATG TATCGCCTAT ACATGGCAGA GGACTGAACA CCTCCTCAGA AGTCAGCGCA
481 GGCCGAGCCT GCTTTGGACA CGGGAGAAAA GAAGCCATTG CTAAGTACTT CAACTGACAG
541 AAACCTTCAC TTGAAAACAA TGATTTTAAAT ATATCTCTTT CTTTTTCTTC CGACATTAGA
601 AACAAAACAA AAAGAACTGT CCTTTCTGCG CTCAAATTTT TCGAGTGTGC CTTTTTATTC
661 ATCTACTTTA TTTTGATGTT TCCTTAATGT GTAATTTACT TATTATAAGC ATGATCTTTT
721 AAAAATATAT TTGGCTTTTA AAG

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GENBANK ID: M14362.1
DEFINITION HUMAN T-CELL SURFACE ANTIGEN CD2 (T11) MRNA, COMPLETE CDS.
VERSION M14362.1 GI:179133
CDS 10..1065
/CODON_START=1

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1 ACCCCTAAGA TGAGCTTTCC ATGTAAATTT GTAGCCAGCT TCCTTCTGAT TTTCAATGTT
61 TCTTCCAAAG GTGCAGTCTC CAAAGAGATT ACGAATGCCT TGGAAACCTG GGGTGCCTTG
121 GGTCAAGACA TCAACTTGGA CATTCTAGT TTTCAAATGA GTGATGATAT TGACGATATA
181 AAATGGGAAA AAACCTCAGA CAAGAAAAAG ATTGCACAA TCAGAAAAGA GAAAGAGACT
241 TTCAAGGAAA AAGATACATA TAAGCTATTT AAAATGGAA CTCTGAAAAT TAAGCATCTG
301 AAGACCGATG ATCAGGATAT CTACAAGGTA TCAATATATG ATACAAAAGG AAAAAATGTG
361 TTGGAAAAAA TATTTGATTT GAAGATTCAA GAGAGGGTCT CAAAACCAA GATCTCCTGG
421 ACTTGTATCA ACACAACCTT GACCTGTGAG GTAATGAATG GAACTGACCC CGAATTAAAC
481 CTGTATCAAG ATGGGAAACA TCTAAAACCT TCTCAGAGGG TCATCACACA CAAGTGGACC
541 ACCAGCCTGA GTGCAAAATT CAAGTGCACA GCAGGGAACA AAGTCAGCAA GGAATCCAGT
601 GTCGAGCCTG TCAGCTGTCC AGAGAAAGGT CTGGACATCT ATCTCATCAT TGGCATATGT
661 GGAGGAGGCA GCCTCTTGAT GGTCTTTGTG GCACTGCTCG TTTTCTATAT CACCAAAGG
721 AAAAAACAGA GGAGTCGGAG AAATGATGAG GAGCTGGAGA CAAGAGCCCA CAGAGTAGCT
781 ACTGAAGAAA GGGGCCGGAA GCCCCACCAA ATTCCAGCTT CAACCCCTCA GAATCCAGCA
841 ACTTCCCAAC ATCCTCCTCC ACCACCTGGT CATCGTTCCC AGGCACCTAG TCATCGTCCC
901 CCGCCTCCTG GACACCGTGT TCAGCACCAG CCTCAGAAGA GGCTCCTGTC TCCGTCGGGC
961 ACACAAGTTC ACCAGCAGAA AGGCCCGCCC CTCCCAGAC CTCGAGTTCA GCCAAAACCT
1021 CCCCATGGGG CAGCAGAAAA CTCATTGTCC CCTTCCTCTA ATTAAAAAAG ATAGAACTG
1081 TATTTTTCAT TAAAAAGCAC TGTGGATTTC TGCCCTCCTG ATGTGCATAT CCGTACTTCC
1141 ATGAGGTGTT TTCTGTGTGC AGAAGATTGT CACCTCCTGA GGCTGTGGGC CACAGCCACC
1201 TCTGCATCTT CGAACTCAGC CATGTGGTAT ACATCTGGAG TTTTGGTCT CCTCAGAGAG
1261 CTCCATCACA CCACTAAGGA GAAGCAATAT AAGTGTGATT GCAAGAATGG TAGAGGACCG
1321 AGCACAGAAA TCTTAGAGAT TTCCTGTCCC CTCTCAGGTC ATGTGTAGAT GCGATAAATC
1381 AAGTGATTGG TGTGCCTGGG TCTCACTACA AGCAGCCTAT CTGCTTAAGA GACTCTGGAG
1441 TTTCTTATGT GCCCTGGTGG ACACTTGCCC ACCATCCTGT GAGTAAAAGT GAAATAAAAG
1501 CTTTGACTAG

GENBANK ID: XM_087746.1DEFINITION HOMO SAPIENS SIMILAR TO KIDNEY AMINOPEPTIDASE M; LEUCINE
ARYLAMINOPEPTIDASE 1 (LOC153726), MRNA.

VERSION XM_087746.1 GI:18561749

CDS 262..639

/CODON_START=1

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1 GAGTTCCATG CCACCTCCCC GCCCTTTACA GACATGCTAT AAGGTCCCCA GCCCAGTCAC
61 TCCGCAGTGC CTCTCTCTTC CTCCCCATGG ACTATACACA GGCCCTGCTT GTCCTGGAGG
121 AAAGTTTGGG CGTCATTATA TAGATCAGGA GACTGAAGTA CTGAAAGGTT AAATGACTTG
181 CCAAAGAATG AGATCTTTTT TTCTAACATT TTACATAATA TCCTCAGAGA AGATCAGGCC
241 CTGGTGACTA GAGCTGTGGC CATGAAGGTG GAAAATTTCA AAACAAGTGA AATACAGGAA
301 CTCTTTGACA TATTTACTTA CAGCAAGGGA GCGTCTATGG CCCGGATGCT TTCTTGTTTC
361 TTGAATGAGC ATTTATTTGT CAGTGCACCT AAGTCATATT TGAAGACATT TTCCTACTCA
421 AACGCTGAGC AAGATGATCT ATGGAGGCAT TTTCAAATGG CCATAGATGA CCAGAGTACA
481 GTTATTTTGC CAGCAACAAT AAAAAACATA ATGGACAGTT GGACACACCA GAGTGGTTTT
541 CCAGTGATCA CTTTAAATGT GTCTACTGGC GTCATGAAAC AGGAGCCATT TTATCTTGAA
601 AACATTAAAA ATCGGACTCT TCTAACCAGC AATAAGTGAC ACATGGATTG TCCCTATTCT
661 TTGGATAAAA AATGGAACCT CACAACCTTT AGTCTGGCTA GA
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GENBANK ID: P31749DEFINITION DICTYOSTELIUM DISCOIDEUM RAC-ALPHA SERINE/THREONINE KINASE HOMOLOG
MRNA, COMPLETE CDS.

VERSION U15210.1 GI:1000068

CDS 1..1335

/CODON_START=1

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61 AGTTGGAAAA AGAGATGGTT CATTCTCAAA GGTGGTGATT TAAGTTATTA TAAAACAAA
121 GGTGAAGTTG TACCATTAGG AGTTATTCAT TTAAATACAT CAGGTCAATAT TAAAAATTCT
181 GATCGTAAGA AAAGAGTTAA TGGATTTGAA GTACAAACAC CATCACGTAC ATATTTCTTA
241 TGTTCAAGAG CAGAGGAAGA ACGTGCAAAA TGGATAGAGA TATTAATTAA TGAAAGAGAA
301 TTATTATTGA ATGGTGGAACA ACAACCAAAG AAATCGGAAA AGGTAGGAGT TGCAGATTTT
361 GAATTATTGA ATTTAGTTGG TAAAGGTAGT TTTGGTAAAG TTATTCAAGT TAGAAAGAAA
421 GATACTGGTG AAGTGTATGC AATGAAAGTT TTATCAAAGA AACATATCGT AGAGCATAAC
481 GAAGTCGAAC ATACATTGAG TGAGCGTAAT ATTCTTCAAA AGATCAATCA CCCATTTTTG
541 GTTAATCTCA ACTACAGTTT TCAAACAGAG GATAAGCTTT ACTTTATCTT GGATTATGTT
601 AATGGTGGTG AGTTATTCTA TCATCTTCAA AAGGACAAA AGTTTACAGA GGATCGTGTC
661 CGTTATTATG GCGCAGAGAT CGTATTGGCA TTGGAACATT TACATTTGTC GGTGTCATC
721 TATAGAGATT TGAAACCAGA GAATTTACTA CTCACCAACG AGGGTCACAT TTGCATGACC
781 GATTTCCGTC TTTGCAAAGA GGGTCTATTG ACACCAACCG ACAAACCTGG TACTTTCTGT
841 GGTACTCCTG AATATTTAGC ACCCGAAGTA CTTCAAGGCA ATGGTTATGG TAAACAAGTG
901 GATTGGTGGA GTTTTGGTTC TCTCCTCTAT GAAATGCTCA CTGGTTTACC ACCATTCTAC
961 AATCAAGACG TCCAAGAGAT GTATCGTAAG ATCATGATGG AGAAATTATC TTTCCACAT
1021 TTCATTTCTC CAGATGCTCG TTCCCTCTTG GAACAACTCT TGGAAAGAGA TCCTGAAAAA
1081 AGACTTGCCG ATCCAAATCT TATTAAAAGA CATCCTTTCT TCCGTTCCAT CGATTGGGAA
1141 CAATTATTCC AAAAGAATAT TCCACCACCA TTCATTCCAA ATGTTAAAGG TTCTGCTGAT
1201 ACCTCTCAAA TTGATCCAGT TTCACTGAT GAAGCTCCTT CTTTAACTAT GGCTGGTGAA
1261 TGTGCTTTAA ATCCGCAACA AAAAAAGAT TTTGAAGGAT TTACATATGT CGCTGAATCT
1321 GAACATTTAA GATAA
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GENBANK ID: NM_000102.2DEFINITION HOMO SAPIENS CYTOCHROME P450, SUBFAMILY XVII (STEROID
17-ALPHA-HYDROXYLASE), ADRENAL HYPERPLASIA (CYP17), MRNA.

VERSION NM_000102.2 GI:13904854

CDS 61..1587

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1 GAGTTGCCAC AGCTCTTCTA CTCCACTGCT GTCTATCTTG CCTGCCGGCA CCCAGCCACC
61 ATGTGGGAGC TCGTGGCTCT CTTGCTGCTT ACCCTAGCTT ATTTGTTTTG GCCCAAGAGA
121 AGGTGCCCTG GTGCCAAGTA CCCCAAGAGC CTCCTGTCCC TGCCCTGCTT GGGCAGCCTG
181 CCATTCCTCC CCAGACATGG CCATATGCAT AACAACCTCT TCAAGCTGCA GAAAAAATAT
241 GGCCCCATCT ATTCTGTTTC TATGGGCACC AAGACTACAG TGATTGTCGG CCACCACCAG
301 CTGGCCAAGG AGGTGCTTAT TAAGAAGGGC AAGGACTTCT CTGGGCGGCC TCAAATGGCA
361 ACTCTAGACA TCGCGTCCAA CAACCGTAAG GGTATCGCCT TCGCTGACTC TGGCGCACAC
421 TGGCAGCTGC ATCGAAGGCT GCGGATGGCC ACCTTTGCCC TGTTCAGGA TGGCGATCAG
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5 481 AAGCTGGAGA AGATCATTTG TCAGGAAATC AGTACATTGT GTGATATGCT GGCCACCCAC
541 AACGGACAGT CCATAGACAT CTCCTTTCCT GTCTTCGTGG CGGTAACCAA TGTCATCTCC
601 TTGATCTGCT TCAATACCTC CTACAAGAAT GGGGACCCTG AGTTGAATGT CATAAGAAT
661 TACAATGAAG GCATCATAGA CAACCTGAGC AAAGACAGCC TGGTGGACCT AGTCCCCTGG
721 TTGAAGATTT TCCCCAACAA AACCTGGAA AAATTAAAGA GCCATGTTAA AATACGAAAT
781 GATCTGCTGA ATAAAATACT TGAAAATTAC AAGGAGAAAT TCCGGAGTGA CTCTATCACC
841 AACATGCTGG ACACACTGAT GCAAGCCAAG ATGAACTCAG ATAATGGCAA TGCTGGCCCA
901 GATCAAGATT CAGAGCTGCT TTCAGATAAC CACATTCTCA CCACCATAGG GGACATCTTT
10 961 GGGGCTGGCG TGGAGACCAC CACCTCTGTG GTTAAATGGA CCCTGGCCTT CCTGCTGCAC
1021 AATCCTCAGG TGAAGAAGAA GCTCTACGAG GAGATTGACC AGAATGTGGG TTTCAGCCGC
1081 ACACCAACTA TCAGTGACCG TAACCGTCTC CTCCTGCTGG AGGCCACCAT CCGAGAGGTG
1141 CTTGCGCTCA GGCCCGTGGC CCCTATGCTC ATCCCCACA AGGCCAACGT TGACTCCAGC
1201 ATCGGTGAGT TTGCTGTGGA CAAGGGCACA GAAGTTATCA TCAATCTGTG GGCGCTGCAT
1261 CACAATGAGA AGGAGTGGCA CCAGCCGGAT CAGTTCATGC CTGAGCGTTT CTTGAATCCA
15 1321 GCGGGGACCC AGCTCATCTC ACCGTCAGTA AGCTATTTGC CCTTCGGAGC AGGACCTCGC
1381 TCCTGTATAG GTGAGATCCT GGCCCGCCAG GAGCTCTTCC TCATCATGGC CTGGCTGCTG
1441 CAGAGGTTCTG ACCTGGAGGT GCCAGATGAT GGCAGCTGC CCTCCCTGGA AGGCATCCCC
1501 AAGGTGGTCT TTCTGATCGA CTCTTTCAAA GTGAAGATCA AGGTGCGCCA GGCTGGAGG
1561 GAAGCCCAGG CTGAGGGTAG CACCTAAAGG CTGTAACCTA CAGCCCCTGT CCACCCATG
20 1621 TGGCCCCACA ACACAGATTT AGAGATACAA CCCCCACCC TTCTCCGCCA TTCTCCCTA
1681 CTCCCAACCC ACTCTGCCTT CTTTTTCAGC TTGTGGCAAT GCCAGTGATG TGCATAACA
1741 GTTTTTTTTT TTTCC

25 GENBANK ID: NM_001662
DEFINITION HOMO SAPIENS ADP-RIBOSYLATION FACTOR 5 (ARF5), MRNA.
VERSION NM_001662.2 GI:6995999
CDS 37..579
/CODON_START=1

30 1 CCGCGTCGGT GCCCGCGCCC CTCCCCGGGC CCGCCATGG GCCTCACCCT GTCCGCGCTC
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121 GGCAAGACCA CAATCCTGTA CAACTGAAG TTGGGGGAGA TTGTCACCAC CATCCCAACC
181 ATAGGCTTCA ATGTAGAAAC AGTGAATAT AAGAACATCT GTTTCACAGT CTGGGACGTG
241 GGAGGCCAGG ACAAGATTCG GCCTCTGTGG CGGCACTACT TCCAGAACAC TCAGGGCCTC
35 301 ATCTTTGTGG TGGACAGTAA TGACCGGGAG CGGGTCCAAG AATCTGCTGA TGAATCCAG
361 AAGATGCTGC AGGAGGACGA GCTGCGGGAT GCAGTGCTGC TGGTATTTGC CAACAAGCAG
421 GACATGCCCA ACGCCATGCC CGTGAGCGAG CTGACTGACA AGCTGGGGCT ACAGCACTTA
481 CGCAGCCGCA CGTGGTATGT CCAGGCCACC TGTGCCACCC AAGGCACAGG TCTGTACGAT
541 GGTCTGGACT GGCTGTCCCA CGAGCTGTCA AAGCGCTAAC CAGCCAGGGG CAGGCCCTG
40 601 ATGCCCGGAA GCTCCTGCGT GCATCCCCGG GATGACCAGA CTCCCGGACT CCTCAGGCAG
661 TGCCCTTTCC TCCCACCTTT CCTCCCCCAT AGCCACAGGC CTCTGCTCCT GCTCCTGCCT
721 GCATGTTCTC TCTGTTGTTG GAGCCTGGAG CCTTGCTCTC TGGGCACAGA GGGGTCCACT
781 CTCCTGCCTG CTGGGACCTA TGGAAAGGGC TTCCTGGCCA AGGCCCCCTC TTCCAGAGGA
841 GGAGCAGGGA TCTGGGTTTC CTTTTTTTTT TCTGTTTTGG GTGTACTCTA GGGGCCAGGT
45 901 TGGGAGGGGG AAGGTGAGGG CTTGGGTGG TGCTATAATG TGGCACTGGA TCTTGAGTAA
961 TAAATTTGCT GTGGTTTG

50 GENBANK ID: XM_048886.3
DEFINITION HOMO SAPIENS MICROSOMAL GLUTATHIONE S-TRANSFERASE 1 (MGST1), MRNA.
VERSION XM_048886.3 GI:18580621
CDS 89..556
/CODON_START=1

55 1 AGTCCCTGCA TTGCGCGCGA CCCGCGGCG GGACAGGCTT GCTGCTTCCT CTCCTCGGC
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121 AGTATTCATG GCTTTTGCAT CCTATGCAAC AATTATTCTT TCAAAAATGA TGCTTATGAG
181 TACTGCAACT GCATTCTATA GATTGACAAG AAAGGTTTTT GCCAATCCAG AAGACTGTGT
241 AGCATTGCGC AAAGGAGAAA ATGCCAAGAA GTATCTTCGA ACAGATGACA GAGTAGAACG
301 TGTACGCAGA GCCACCTGA ATGACCTTGA AAATATTATT CCATTTCTTG GAATTGGCCT
60 361 CCTGTATTCC TTGAGTGGTC CCGACCCCTC TACAGCCATC CTGCACTTCA GACTATTTGT
421 CGGAGCACGG ATCTACCACA CCATTGCATA TTTGACACCC CTTCCCCAGC CAAATAGAGC
481 TTTGAGTTT TTTGTTGGAT ATGGAGTTAC TCTTTCCATG GCTTACAGGT TGCTGAAAAG
541 TAAATTGTAC CTGTAAAGAA AATCATACAA CTCAGCATCC AGTTGGCTTT TTAAGAATTCT
601 TGTACTTCCA ATTTATAATG AATACTTTCT TAGATTTTAG GTAGGAGGGG AGCAGAGGAA
35 661 TTATGAACTG GGGTAAACCC ATTTTGAATA TTAGCATTGC CAATATCCTG TATTCTTGT
721 TTACATTTGG ATTAGAAATT TAACATAGTA ATTCTTAAGT CTTTTGTCTG ATTTTAAAG

781 TACTTTCTTA TAAATTGGA TCATGTTATG ATTTGTAACA TTCACACAAC ACCTCACTTT
841 TGAATCTATA AAAGAATTGC ACGTATGAGA AACCTATATT TCAATACTGC TGAAACAGAC
901 ATGAAATAAA GAATTTAAAG AATG

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GENBANK ID: X02162

DEFINITION HUMAN MRNA FOR APOLIPOPROTEIN AI (APO AI)=.

VERSION X02162.1 GI:28771

CDS 87..890

/CODON_START=1

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61 AAGGAGGTCC CCCACGGCCC TTCAGGATGA AAGCTGCGGT GCTGACCTTG GCCGTGCTCT

121 TCCTGACGGG GAGCCAGGCT CGGCATTCTT GGCAGCAAGA TGAACCCCCC CAGAGCCCCCT

181 GGGATCGAGT GAAGGACCTG GCCACTGTGT ACGTGGATGT GCTCAAAGAC AGCGGCAGAG

241 ACTATGTGTC CCAGTTTGAA GGCTCCGCCT TGGGAAAACA GCTAAACCTA AAGCTCCTTG

15

301 ACAACTGGGA CAGCGTGACC TCCACCTTCA GCAAGCTGCG CGAACAGCTC GGCCCTGTGA

361 CCCAGGAGTT CTGGGATAAC CTGGAAGAGG AGACAGAGGG CCTGAGGCAG GAGATGAGCA

421 AGGATCTGGA GGAGGTGAAG GCCAAGGTGC AGCCCTACCT GGACGACTTC CAGAAGAAGT

481 GGCAGGAGGA GATGGAGCTC TACCGCCAGA AGGTGGAGCC GCTGCGCGCA GAGCTCCAAG

541 AGGGCGCGCG CCAGAAGCTG CACGAGCTGC AAGAGAAGCT GAGCCCACTG GGCGAGGAGA

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601 TCGCGGACCG CGCGCGCGCC CATGTGGACG CGCTGCGCAC GCATCTGGCC CCTACAGCG

661 ACGAGCTGCG CCAGCGCTTG GCCGCGCGCC TTGAGGCTCT CAAGGAGAAC GGCGGCGCCA

721 GACTGGCCGA GTACCACGCC AAGGCCACCG AGCATCTGAG CACGCTCAGC GAGAAGGCCA

781 AGCCCGCGCT CGAGGACCTC CGCCAAGGCC TGCTGCCCGT GCTGGAGAGC TTCAAGGTCA

841 GCTTCCTGAG CGCTCTCGAG GAGTACACTA AGAAGCTCAA CACCCAGTGA GGCGCCCGCC

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901 GCCGCCCGCC TTCCCGGTGC TCAGAATAAA CGTTTCCAAA GTGGGAAAAA AAAAAAAG

961 AATTC

GENBANK ID: XM_007441.1

DEFINITION HOMO SAPIENS PRESENILIN 1 (ALZHEIMER DISEASE 3) (PSEN1), MRNA.

VERSION XM_007441.1 GI:11435041

CDS 249..1652

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1 TGGGACAGGC AGCTCCGGGG TCCGCGGTTT CACATCGGAA ACAAAACAGC GGCTGGTCTG

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121 CTGGGAGCCT GCAAGTGACA ACAGCCTTTG CGGTCCCTTAG ACAGCTTGGC CTGGAGGAGA

181 ACACATGAAA GAAAGAACCT CAAGAGGCTT TGTTTTCTGT GAAACAGTAT TTCTATACAG

241 TTGCTCCAAT GACAGAGTTA CCTGCACCGT TGTCCTACTT CCAGAATGCA CAGATGTCTG

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301 AGGACAACCA CCTGAGCAAT ACTGTACGTA GCCAGAATGA CAATAGAGAA CGGCAGGAGC

361 ACAACGACAG ACGGAGCCTT GGCCACCCTG AGCCATTATC TAATGGACGA CCCCAGGGTA

421 ACTCCCGGCA GGTGGTGGAG CAAGATGAGG AAGAAGATGA GGAGCTGACA TTGAAATATG

481 GCGCCAAGCA TGTGATCATG CTCTTTGTCC CTGTGACTCT CTGCATGGTG GTGGTCGTGG

541 CTACCATTAA GTCAGTCAGC TTTTATACCC GGAAGGATGG GCAGCTAATC TATACCCCAT

601 TCACAGAAGA TACCGAGACT GTGGGCCAGA GAGCCCTGCA CTCAATTCTG AATGCTGCCA

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661 TCATGATCAG TGTCATTGTT GTCATGACTA TCCTCCTGGT GGTTCCTGTAT AAATACAGGT

721 GCTATAAGGT CATCCATGCC TGGCTTATTA TATCATCTCT ATTGTTGCTG TTCTTTTTTT

781 CATTCAATTA CTTGGGGGAA GTGTTTAAAA CCTATAACGT TGCTGTGGAC TACATTACTG

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901 CACTTCGACT CCAGCAGGCA TATCTCATTA TGATTAGTGC CCTCATGGCC CTGGTGTTTA

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961 TCAAGTACCT CCCTGAATGG ACTGCGTGGC TCATCTTGGC TGTGATTTCA GTATATGATT

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1081 GAAATGAAAC GCTTTTCCA GCTCTCATTT ACTCCTCAAC AATGGTGTGG TTGGTGAATA

1141 TGGCAGAAGG AGACCCGGA GCTCAAAGGA GAGTATCCAA AAATTCCAAG TATAATGCAG

1201 AAAGCACAGA AAGGGAGTCA CAAGACACTG TTGCAGAGAA TGATGATGGC GGGTTCAGTG

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1261 AGGAATGGGA AGCCCAGAGG GACAGTCATC TAGGGCCTCA TCGCTCTACA CCTGAGTCAC

1321 GAGCTGCTGT CCAGGAACTT TCCAGCAGTA TCCTCGCTGG TGAAGACCCA GAGGAAAGGG

1381 GAGTAAACT TGGATTGGGA GATTTCATTT TCTACAGTGT TCTGGTTGGT AAAGCCTCAG

1441 CAACAGCCAG TGGAGACTGG AACACAACCA TAGCCTGTTT CGTAGCCATA TTAATTGGTT

1501 TGTGCCTTAC ATTATTACTC CTTGCCATTT TCAAGAAAGC ATTGCCAGCT CTTCCAATCT

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1561 CCATCACCTT TGGGCTTGT TTTACTTTT CCACAGATTA TCTTGTACAG CTTTTTATGG

1621 ACCAATTAGC ATTCCATCAA TTTTATATCT AGCATATTTG CGGTTAGAAT CCCATGGATG

1681 TTTCTTCTTT GACTATAACA AAATCTGGGG AGGACAAAGG TGATTTTCCT GTGTCCACAT

1741 CTAACAAAGT CAAGATTCCC GGCTGGACTT TTGCAGCTTC CTTCCAAGTC TTCTGACCA

1801 CTTGCACTA TTGGACTTTG GAAGGAGGTG CCTATAGAAA ACGATTTTGA ACATACTTCA

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1861 TCGCAGTGGA CTGTGTCCCT CGGTGCAGAA ACTACCAGAT TTGAGGGACG AGGTCAAGGA

1921 GATATGATAG GCCCGGAAGT TGCTGTGCCC CATCAGCAGC TTGACGCGTG GTCACAGGAC

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1981 GATTTCACTG ACACTGCGAA CTCTCAGGAC TACCGTTACC AAGAGGTTAG GTGAAGTGGT
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2101 TATTAACCTGA ATTCTGAACT TTTCAGGAGG TACTGTGAGG AAGAGCAGGC ACCAGCAGCA
2161 GAATGGGGAA TGGAGAGGTG GGCAGGGGTT CCAGCTTCCC TTTGATTTT TGCTGCAGAC
2221 TCATCCTTTT TAAATGAGAC TTGTTTTCCC CTCTCTTTGA GTCAAGTCAA ATATGTAGAT
2281 TGCCTTTGGC AATTCTTCTT CTCAAGCACT GACACTCATT ACCGTCTGTG ATTGCCATTT
2341 CTTCCCAAGG CCAGTCTGAA CCTGAGGTTG CTTTATCCTA AAAGTTTTAA CCTCAGGTTT
2401 CAAATTCAGT AAATTTTGGG AACAGTACAG CTATTTCTCA TCAATTCTCT ATCATGTTGA
2461 AGTCAAATTT GGATTTTCCA CCAAATTCTG AATTTGTAGA CATACTTGTA CGCTCACTTG
2521 CCCCAGATGC CTCCTCTGTC CTCATTCTTC TCTCCACAC AAGCAGTCTT TTTCTACAGC
2581 CAGTAAGGCA GCTCTGTCGT GGTAGCAGAT GGTCCCATTA TTCTAGGGTC TTACTCTTTG
2641 TATGATGAAA AGAATGTGTT ATGAATCGGT GCTGTCAGCC CTGCTGTCAG ACCTTCTTCC
2701 ACAGCAAATG AGATGTATGC CCAAAGACGG TAGAATTAAA GAAGAGTAAA ATGGCTGTTG
2761 AAGC

GENBANK ID: XM_087242.1
DEFINITION HOMO SAPIENS ARGINYL AMINOPEPTIDASE (AMINOPEPTIDASE B)-LIKE
1(RNPEPL1), MRNA.
VERSION XM_087242.1 GI:18600482
CDS 700..1764
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121 TGGCTGGACC CAGAGCTGAC CTATGGCTGC GCCAAGCCCT TCGTCTTCAC CCAGGGCCAC
181 TCCGTGTGCA ACCGCTCCTT CTTCCCGTGC TTCGACACAC CTGCCGTGAA GTGCACCTAC
241 TCTGCCGTGC TCAAGGCGCC ATCGGGGGTG CAGGTGCTGA TGAGTGCCAC CCGGAGTGCA
301 TACATGGAGG AAGAAGGCGT CTTCCACTTC CACATGGAGC ACCCCGTGCC CGCCTACCTC
361 GTGGCCCTGG TGGCCGGAGA CCTCAAGCCG GCAGACATCG GGCCAGGAG CCGCGTGTGG
421 GCCGAGCCAT GCCTCCTGCC CACGGCCACC AGCAAGCTGT CGGGCGCAGT GGAGCAGTGG
481 CTGAGTGCAG CTGAGCGGCT GTATGGGCCC TACATGTGGG GCAGGTACGA CATGTCTTTC
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601 TCCATCCTGG AGAGCGATGA GTTCTTGGTG ATCGATGTCA TCCACGAGGT GGCCACAGT
661 TGGTTCGGCA CCCGCTGCAC CAAGCCACAG TGGGAAGAGA TGTGGCTGAG CGAGGGCCTG
721 GCCACCTATG ACCAGCGCCG TATCACCACC GAGACCTACG GTGCTGCCTT CACCTGCCTG
781 GAGACTGCCT TCCGCTGGA CGCCCTGCAC CGGCAGATGA AGCTTCTGGG AGAGGACAGC
841 CCGGTCAGCA AACTGCAGGT CAAGCTGGAG CCAGGAGTGA ATCCCAGCCA CCTGATGAAC
901 CTGTTACCT ACAGAAAGGG CTACTGCTTC GTGTACTACC TGTCCAGCT CTGCGGAGAC
961 CCACAGCGCT TTGATGACTT TCTCCGAGCC TATGTGGAGA AGTACAAGTT CACCAGCGTG
1021 GTGGCCCAGG ACCTGCTGGA CTCCTTCTTG AGCTTCTTCC CGGAGCTGAA GGAGCAGAGC
1081 GTGGACTGCC GGGCAGGGCT GGAATTCGAG CGCTGGCTCA ATGCCACAGG CCCGCCGCTG
1141 GCTGAGCCGG ACCTGTCTCA GGGATCCAGC CTGACCCGGC CCGTGGAGGC CCTTTTCCAG
1201 CTGTGGACCG CAGAACCTCT GGACCAGGCA GCTGCCTCGG CCAGCGCCAT TGACATCTCC
1261 AAGTGGAGGA CCTTCCAGAC AGCACTCTTC CTGGACCGGC TCCTGGATGG GTCCCCGCTG
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1441 CACAGGGTGC GCGCTTCTT GGAGAGCCAG ATGTCACGCA TGTACACCAT CCCGCTGTAC
1501 GAGGACCTCT GCACCGGTGC CCTCAAGTCC TTCGCGCTGG AGGTCTTCTA CCAGACGCAG
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1861 CTCTGCCCAG GCCCACAAGC CCTCCCCTG GGCTCTCCCA GGCAGGGAGA ATGGGGAGAG
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1981 CACTGCAGGC CCTGGGGCCA GCGGACAC ACCATGCCTC CTGTCTCAAC ACTGACAGCT
2041 GTGCCTAGCC CCGGATGCCA GCACCTGCCA GGTGCCGCCC CGGGGCAAGG GCGCCAGCAG
2101 CCCTATGGTG ACCGCCACAC TGTGCTTAA GTCTGCCCAG GGGCCAGGC TGTGCTGTCC
2161 CTGCAGCACG CTTCTTGA GGGATCTGAG CCACCTCCC CGCACAGCCC TGCACCCCGC
2221 CCCTAGGGTT GGCAGCCTCA GTTGCCCCCT GGCAGAGGAA CAAGGACACA GACATTCCCT
2281 CAGTGTGGGG GGCAGGGGAC ACAGGGAGAG GATGGTTGTC CCTGGGGAGG GCCCTCTGGC
2341 CCCAGGCAAC CTTAGCCCTC CAGAACAGG AGTCCCAGGA CCCAGGGAGA GTGTGGGGAC
2401 AGGACAGCCT GTCTCTTGA GCTTCTTGG GTGGGAGGCA CAGGGGCAA GCAATACCCC
2461 AGGGAAAGTG GGAGGTGGTG CTGGTGTCT CTCCAGGCC ACCATGCTGG GAGAGGCGGC
2521 CAGAGCCTGG GGCCTCCAGC CTGGGACTGC TGTGATGGGG TATCACGGTG ATGGTCCCAT
2581 TAAACTTCCA CTCTGCAAAC CTG

GENBANK ID: S90469

DEFINITION CYTOCHROME P450 REDUCTASE [HUMAN, PLACENTA, MRNA PARTIAL, 2403 NT].

VERSION S90469.1 GI:247306

CDS 1..2031

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61 CTTTTCAGCA TGACGGACAT GATTCTGTTT TCGCTCATCG TGGGTCTCCT AACCTACTGG
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241 GTGTTCTACG GCTCCAGAC GGGGACTGCA GAGGAGTTTG CCAACCGCCT GTCCAAGGAC
301 GCCCACCGCT ACGGGATGCG AGGCATGTCA GCGGACCCTG AGGAGTATGA CCTGGCCGAC
361 CTGAGCAGCC TGCCAGAGAT CGACAACGCC CTGGTGGTTT TCTGCATGGC CACCTACGGT
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661 CAGTTCTGGC CGGCCGTGTG TGAACACTTT GGGGTGGAAG CCACTGGCGA GGAGTCCAGC
721 ATTGCGCAGT ACGAGCTTGT GGTCCACACC GACATAGATG CGGCCAAGGT GTACATGGGG
781 GAGATGGGCC GGCTGAAGAG CTACGAGAAC CAGAAGCCCC CTTTGTATGC CAAGAATCCG
841 TTCCTGGCTG CAGTCACCAC CAACCGGAAG CTGAACCAGG GAACCGAGCG CCACCTCATG
901 CACCTGGAAT TGGACATCTC GGACTCCAAA ATCAGGTATG AATCTGGGGA CCACGTGGCT
961 GTGTACCCAG CCAACGACTC TGCTCTCGTC AACCAGCTGG GCAAAATCCT GGGTGCCGAC
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1621 GGCTTCATCC AGGAGCGGGC CTGGCTGCGA CAGCAGGGCA AGGAGGTGGG GGAGACGCTG
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1801 AAGGTCTACG TCCAGCACCT GCTAAAGCAA GACCGAGAGC ACCTGTGGAA GTTGATCGAA
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1921 ACCTTCTACG ACATCGTGGC TGAGCTCGGG GCCATGGAGC ACGCGCAGGC GGTGGACTAC
1981 ATCAAGAAAC TGATGACCAA GGGCCGCTAC TCCCTGGACG TGTGGAGCTA GGGGCCTGCC
2041 TGCCCCACCC ACCCCACAGA CTCCGGCCTG TAATCAGCTC TCCTGGCTCC CTCCCGTAGT
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2281 GGTGGCTGCA CAGAAGGGCT CTTTCTCTCT GCTGAGCTGG CCCAGCCCCC CCACGTGATT
2341 TCCAGTGAGT GTAAATAATT TTAAATAACC TCTGGCCCTT GGAATAAAGT TCTGTTTTCT
2401 GTA
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GENBANK ID: NM_006254.1

DEFINITION HOMO SAPIENS PROTEIN KINASE C, DELTA (PRKCD), MRNA.

VERSION NM_006254.1 GI:5453969

CDS 59..2089

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181 GAAACACTG GTGCAGAAGA AGCCGACCAT GTATCCTGAG TGGAAGTCGA CGTTCGATGC
241 CCACATCTAT GAGGGGCGCG TCATCCAGAT TGTGCTAATG CGGGCAGCAG AGGAGCCAGT
301 GTCTGAGGTG ACCGTGGGTG TGTCGGTGCT GGCCGAGCGC TGCAAGAAGA ACAATGGCAA
361 GGCTGAGTTC TGGCTGGACC TGCAGCCTCA GGCCAAGGTG TTGATGTCTG TTCAGTATTT
421 CCTGGAGGAC GTGGATTGCA AACAATCTAT GCGCAGTGAG GACGAGGCCA AGTTCCCAAC
481 GATGAACCGC CGCGGAGCCA TCAAACAGGC CAAAATCCAC TACATCAAGA ACCATGAGTT
541 TATCGCCACC TTCTTTGGGC AACCACCTT CTGTTCTGTG TGCAAAGACT TTGTCTGGGG
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5 661 CGACAAGATC ATCGGCAGAT GCACTGGCAC CGCGGCCAAC AGCCGGGACA CTATATTCCA
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841 TGAAGACTGC GGCATGAATG TGCACCATAA ATGCCGGGAG AAGGTGGCCA ACCTCTGCGG
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961 ATCAGACTCA GCCTCCTCAG AGCCTGTTGG GATATATCAG GGTTTCGAGA AGAAGACCGG
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1141 GAAGGTGCTG CTTGGAGAGC TGAAGGGCAG AGGAGAGTAC TCTGCCATCA AGGCCCTCAA
1201 GAAGGATGTG GTCCTGATCG ACGACGACGT GGAGTGCACC ATGGTTGAGA AGCGGGTGCT
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1381 CAAAGGCCGC TTTGAACTCT ACCGTGCCAC GTTTTATGCC GCTGAGATAA TGTGTGGACT
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1921 GCCCAAAGTG AAGTCACCCA GAGACTACAG TAACTTTGAC CAGGAGTTCC TGAACGAGAA
1981 GGCGCGCCTC TCCTACAGCG ACAAGAACCT CATCGACTCC ATGGACCAGT CTGCATTGCG
2041 TGGCTTCTCC TTTGTGAACC CCAAATTCGA GCACCTCCTG GAAGATTGAG GTTCCTGGAC
2101 AGAT

GENBANK ID: X61971.1

DEFINITION H.SAPIENS MRNA FOR MACROPAIN SUBUNIT DELTA.

VERSION X61971.1 GI:296733

CDS <1..543

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121 CACAGCATTG AACTGAATGA GCCTCCACTG GTCCACACAG CAGCCAGCCT CTTTAAGGAG
181 ATGTGTTACC GATACCGGGA AGACCTGATG GCGGGAATCA TCATCGCAGG CTGGGACCCT
241 CAAGAAGGAG GGCAGGTCTA CTCAGTGCCG ATGGGGGGTA TGATGGTAAG GCAGTCCTTT
301 GCCATTGGAG GCTCCGGGAG CTCCTACATC TATGGCTATG TTGATGCTAC CTACCGGGAA
361 GGCATGACCA AGGAAGAGTG TCTGCAATTC ACGGCCAATG CTCTCGCTTT GGCCATGGAG
40 421 CGGGATGGCT CAGTGGAGG AGTGATCCGC CTGGCAGCCA TTGCAGAGTC AGGGGTAGAG
481 CGGCAAGTAC TTTTGGGAGA CCAGATACCC AAATTCGCCG TTGCCACTTT ACCACCGGCC
541 TGAATCCTGG GATTCTAGTA TGCAATAAGA GATGCCCTGT ACTGATGCAA AATTTAATAA
601 AGTTTGTGTC AGAGAAAAAA AAAA

45 GENBANK ID: AH005909.1

GENBANK ID: XM_088424.1

DEFINITION HOMO SAPIENS RETINOID X RECEPTOR, ALPHA (RXRA), MRNA.

VERSION XM_088424.1 GI:18571706

CDS 519..1016

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121 TTGAAAAGGT GATGTGTGGG GAGTGCGGCT CATCACTGAG TAGAGAGGTA GAATTTCTAT
181 TTAACCAGAC CTGTAGTAGT ATTACCAATC CAGTTCAATT AAGGTGATTT TTTGTAATTA
241 TTATTATTTT GGTGGGACAA TCTTTAATTT TCTAAAGATA GCACTAACAT CAGCTCATTA
301 GCCACCTGTG CTTGTCCCCG CCTTGGCCCCG GCTGGATGAA GCGGCTTCCC CGCAGGGCCC
361 CCACTTCCCA GTGGCTGCTT CCTGGGGACC CAGGGCACCC CGGCACCTTC AGGCACGCTC
60 421 CTCAGCTGGT CACCTCCCGG CTTTGCCGTT CAGATGGGGC TCCTGAGGCT CAGGAGTGAA
481 GATGCCACAG AGCCGGGCTC CCCTAGGCTG CGTCGGGCAT GCTTGGGAAGC TGGCCTGCCA
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601 CCTCTGGGAA GGACAGCCCT GACCTTCGGT TTTCCGAGCA CCGTGTTCCT CAAGAATTCT
661 GGGCTTGCCG CCTGGTGGCA GTGCTGGAGA TGACCCCGAG CCCCTCCCCG TGGGGCACCC
65 721 AGGAGGGCCC TGCCGAAATG TGCAGCCTGT GGGTAGTCGG CTGGTGTCCC TGTCGTGGAG
781 CTGGGGTGCG TGATCTGGTG CTCGTCCACG CAGGTGTGTG GTGTAAACAT GTATGTGCTG

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841 TACAGAGAGA CGCGTGTGGA GAGAGCCGCA CACCAGCGCC ACCCAGGAAA GGCGGAGCGG
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961 CTTCGTGTAA GCAAGTACAT AAGGACCCCTC CTTTGGTGAA ATCCGGGTTC GAATGAATAT
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1201 AATCTATTTT TGTACAAATG TAATTTTATC CCTCATGTAT ACTTGATAT GGCGGGGGGA
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1321 GCCAGGACAG ACGATGGCAG AGGAGAGGGC TCCTGTGACG GCGGCGAGGC TTGGGAGGAA
1381 ACCGCCGCAA TGGGGGTGTC TTCCCTCGGG GCAGGAGGGT GGGCCTGAGG CTTTCAAGGG
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1561 GCCGCAGGTG CCAGGGTTTG ATGGACAGTA GCATTAGAAAT TGTGGAAGAG GAACACGCAA
1621 AGGGAGAAGT GTGAGAGGAG AAACAAAATA TGAGCGTTTA AAATACATCG CCATTTCAG

GENBANK ID: U41745.1

DEFINITION HUMAN PDGF ASSOCIATED PROTEIN MRNA, COMPLETE CDS.

VERSION U41745.1 GI:1136583

CDS 22..567

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121 GCCAGGGAAG AAGAGGAGCA AAAAGAAGGT GGAGATGGGG CTGCAGGTGA CCCCCAAAAG
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241 CGCAAAGGCG TTGAAGGGCT CATCGACATC GAGAACCCCA ACCGGGTGGC ACAGACAACC
301 AAAAAGGTCA CACAACCTGA TCTGGACGGG CCAAAGGAGC TTTCGAGGAG AGAACGAGAA
361 GAGATTGAGA AGCAGAAGGC AAAAGAGCGT TACATGAAAA TGCACTTGGC CGGGAAGACA
421 GAGCAAGCCA AGGCTGACCT GGCCCGGCTG GCCATCATCC GGAAACAGCG GGAGGAGGCT
481 GCGCGGAAGA AGGAAGAGGA AAGGAACGAT AAAGACGATG CCACATTGTC AGGAAAACGA
541 ATGCAGTCAC TCTCCCTGAA TAAGTAACTG CGACCCGTGG GAGGAGATGC CGGGGACCTG
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661 CAGCCCTCA TGGCCAGGAG CCCCCATGC CTGGGCTCC TCTTCATCTT GGCACAGAAA
721 TTGTTTGGGG GATGGGGGGG GGACTGGGGG AGGGGTAGCT GCTATCTTTG AGACAGAAAG
781 ATGCAGGACA GCATTTTCATA TGTAACCATT TGAATGTTTT TGCTGTTTTT AGAATTC

GENBANK ID: AH002617.1

GENBANK ID: XM_034862.1

DEFINITION HOMO SAPIENS INTERFERON REGULATORY FACTOR 1 (IRF1), MRNA.

VERSION XM_034862.1 GI:14726087

CDS 197..1174

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301 GATCCCATGG AAGCATGCTG CCAAGCATGG CTGGGACATC AACAAGGATG CCTGTTTGT
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421 GACGTGGAAG GCCAACTTTC GCTGTGCCAT GAACTCCCTG CCAGATATCG AGGAGGTGAA
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781 GCACATCCCA GTGGAAGTTG TGCCGGACAG CACCAGTGAT CTGTACAACT TCCAGGTGTC
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1981 ACACATAGGA CGTGTGTAAA TATGTACATT TGTCTTTTA TAAAAGTAA ATTGTT

GENBANK ID: AJ310549.1

DEFINITION HOMO SAPIENS MRNA FOR CLP-36 PROTEIN.

VERSION AJ310549.1 GI:13160404

CDS 1..990

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301 CCATACAAGA TGAATTTAGC CTCTGAACCC CAGGAGGTCC TGCACATAGG AAGCGCCAC
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661 GAAATCCTGG AGTCTGAAGA AAAAGGGGAT CCCAACAAGC CCTCAGGATT CAGAAGTGTT
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841 CCTGAGTGTT ATGTGTGCAC TGACTGTGGC ACCAACCTGA AACAGAAGGG CCATTTCTTT
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961 TATGAAGTGG TCACTGTGTT CCCCAGTGA GCCAGCAGAT CTGACCACTG TTCTCCAGCA
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1081 GCTTACTTTG GTT

GENBANK ID: XM_016642.3

DEFINITION HOMO SAPIENS ADENYLATE KINASE 3 (AK3), MRNA.

VERSION XM_016642.3 GI:16163712

CDS 145..816

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121 ACCGCCCCC TCCTCGCGAA GGCAATGGCT TCCAAACTCC TGCGCGCGGT CATCCTCGGG
181 CCGCCCGGCT CGGGCAAGGG CACCGTGTGC CAGAGGATCG CCCAGAACTT TGGTCTCCAG
241 CATCTCTCCA GCGGCCACTT CTTGCGGGAG AACATCAAGG CCAGCACCAG AGTTGGTGAG
301 GTGGCAAAGC AGTATATAGA GAAAAGTCTT TTGGTTCCAG ACCATGTGAT CACACGCCTA
361 ATGATGTCCG AGTTGGAGAA TAGGCGTGGC CAGCACTGGC TCCTTGATGG TTTTCTTAGG
421 ACATTAGGAC AAGCCGAGGC CCTGGACAAA ATCTGTGAAG TGGATCTAGT GATCAGTTTG
481 AATATTCCAT TTGAAACACT TAAAGATCGT CTCAGCCGCC GTTGGATTCA CCCTCCTAGC
541 GGAAGGGTAT ATAACCTGGA CTTCATCCA CCTCATGTAC ATGGTATTGA TGACGTCCT
601 GGTGAACCAT TAGTCCAGCA GGAGGATGAT AAACCCGAAG CAGTTGCTGC CAGGCTAAGA
661 CAGTACAAAG ACGCGGCAAA GCCAGTCATT GAATTATACA AGAGCCGAGG AGTGCTCCAC
721 CAATTTCCG GAACGGAGAC GAACAAAATC TGGCCCTACG TTTACACACT TTTCTCAAAC
781 AAGATCACAC GTATTCAGTC CAAAGAAGCA TATTGACCCT GCCCAATGGA AGAACCAGGA
841 AGATGTGGTC ATTCATTCAA TAGTGTGTGT AGTATTGGTG CTGTGTCCAA ATTAGAAGCT
901 AGCTGAGGTA GCTTGCAGCA TCTTTTCTAG TTGAAATGGT GAACTGATAG GAAAACAAAT
961 GAGTAGAAAG AGTTCATGAA GAGGCCCTCC TCTGCCTTTC AAAAGGGTGG TCACCTACAC
1021 ATGTTTAAGG TGTCTCTGCA CATGTCTCAA GCCCATCACA AGAAAGCAAG TACAGTGTGG

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1081 ATTTCAAATG GTGTGTAAC TCAGCTCCAG CTGGTTTTTG ACAGCTGTTG CTGTGGTAAT
1141 ATTTTTTACA TGTGATGGTG ATAGTCTCTG GTTCTCCCCA TCCCCACAAA GGCTGTTGAA
1201 CCACAGCACC AGGAAGCCTG AGAATGAATC CTGAGGGCTC TAGCCCAGGC TTTGTCCCAG
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1381 TTTTCTACAT CCACACTCCA TAGAGTCTCT CCTTTTCAGA TATCCTGGGA TGAAAGAATT
1441 TGGCTTTTTT TTTTTTTTTT TTTTGTACAT CTGTTTTTAC TCTTAGGCTT TTAAACAATA
1501 GTTATTGCTC TTATCCCTCT CAGATTCTAA TAACTGAGAG TGATGGGGCT ATATTGAATC
1561 TCTGTATGCA CTGAGAAGTG AGCTATGAAG AGGATCTTAT TAACTGCTG GTCTGACTTT
1621 ATGGATTGAC ACTGTTCCCT TCTTTTATTG TG

GENBANK ID: Z35307.1

DEFINITION H.SAPIENS MRNA FOR ENDOTHELIN-CONVERTING-ENZYME 1.

VERSION 235307.1 GI:535181

CDS 38..2299

/CODON_START=1

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1 CGCCCCCCCC GTGTCCGCCC TGCTGTCCGC GCTGGGGATG TCGACGTACA AGCGGGCCAC
61 GCTGGACGAG GAGGACCTGG TGGACTCGCT CTCCGAGGGC GACGCATACC CCAACGGCCT
121 GCAGGTGAAC TTCCACAGCC CCCGGAGTGG CCAGAGGTGC TGGGCTGCAC GGACCCAGGT
181 GGAGAAGCGG CTGGTGGTGT TGGTGGTACT TCTGGCGGCA GGACTGGTGG CCTGCTTGGC
241 AGCACTGGGC ATCCAGTACC AGACAAGATC CCCCTCTGTG TGCCTGAGCG AAGCTTGTGT
301 CTCAGTGACC AGCTCCATCT TGAGCTCCAT GGACCCACCA GTGGACCCCT GCCATGACTT
361 CTTCAGCTAC GCCTGTGGGG GCTGGATCAA GGCCAACCCA GTCCCTGATG GCCACTCACG
421 CTGGGGGACC TTCAGCAACC TCTGGGAACA CAACCAAGCA ATCATCAAGC ACCTCCTCGA
481 AAACCTCACG GCCAGCGTGA GCGAGGCAGA GAGAAAGGCG CAAGTATACT ACCGTGCGTG
541 CATGAACGAG ACCAGGATCG AGGAGCTCAG GGCCAACCTT CTAATGGAGT TGATTGAGAG
601 GCTCGGGGGC TGGAACATCA CAGGTCCCTG GGCCAAGGAC AACTTCCAGG ACACCCTGCA
661 GGTGGTCACC GCCCACTACC GCACCTCACC CTTCTTCTCT GTCTATGTCA GTGCCGATTC
721 CAAGAACTCC AACAGCAACG TGATCCAGGT GGACCACTCT GGCTTGGGCT TGCCCTCGAG
781 AGACTATTAC CTGAACAAA CTGAAAACGA GAAGGTGCTG ACCGGATATC TGAACATACAT
841 GGTCCAGCTG GGAAGCTGC TGGGCGGCGG GGACGAGGAG GCCATCCGGC CCCAGATGCA
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961 TGATGAGGAG CTCACTTACC ACAAGTGAC GTGCAGCCGAG CTGCAGACCT TGGCACCCGC
1021 CATCAACTGG TTGCCTTTTC TCAACACCAC CTTCTACCCC GTGGAGATCA ATGAATCCGA
1081 GCCTATTGTG GTCTATGACA AGGAATACCT TGAGCAGATC TCCACTCTCA TCAACACCAC
1141 CGACAGATGC CTGCTCAACA ACTACATGAT CTGGAACCTG GTGCGGAAAA CAAGCTCCTT
1201 CCTTGACCAG CGCTTTCAGG ACGCCGATGA GAAGTTCATG GAAGTCATGT ACGGGACCAA
1261 GAAGACCTGT CTTCTCGCTT GGAAGTTTTG CGTGAGTGAC ACAGAAAACA ACCTGGGCTT
1321 TGCGTTGGGC CCCATGTTTG TCAAAGCAAC CTTGCGCGAG GACAGCAAGA GCATAGCCAC
1381 CGAGATCATC CTGGAGATTA AGAAGGCATT TGAGGAAAGC CTGAGCACCC TGAAGTGGAT
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1501 ATACCCCAAC TTCATCATGG ATCCCAAGGA GCTGGACAAA GTGTTTAATG ACTACACTGC
1561 AGTTCCAGAC CTCTACTTTG AAAATGCCAT GCGGTTTTTC AACTTCTCAT GGAGGGTCAC
1621 TGCCGATCAG CTCAGGAAAG CCCCCAACAG AGATCAGTGG AGCATGACCC CGCCCATGGT
1681 GAACGCCTAC TACTCGCCCA CCAAGAATGA GATTGTGTTT CCGGCCGGGA TCCTGCAGGC
1741 ACCATTCTAC ACACGCTCCT CACCCAAGGC CTTAAACTTT GGTGGCATAG GTGTCGTCGT
1801 GGGCCATGAG CTGACTCATG CTTTTGATGA TCAAGGACGG GAGTATGACA AGGACGGGAA
1861 CCTCCGGCCA TGGTGGAGA ACTCATCCGT GGAGGCCTTC AAGCGTCAGA CCGAGTGCAT
1921 GGTAGAGCAG TACAGCAACT ACAGCGTGAA CGGGGAGCCG GTGAACGGGC GGCACACCCT
1981 GGGGGAGAAC ATCGCCGACA ACGGGGGTCT CAAGGCGGCC TATCGGGCTT ACCAGAACTG
2041 GGTGAAGAAG AACGGGGCTG AGCACTCGCT CCCCACCCTG GGCTCACCA ATAACCAGCT
2101 CTTCTTCCTG GGCTTTGCAC AGGTCTGGTG CTCCGTCCGC ACACCTGAGA GCTCCACGA
2161 AGGCCTCATC ACCGATCCCC ACAGCCCCCTC TCGCTTCCGG GTCATCGGCT CCCTCTCCAA
2221 TTCCAAGGAG TTCTCAGAAC ACTTCCGCTG CCCACCTGGC TCACCCATGA ACCCGCCTCA
2281 CAAGTGCGAA GTCTGGTAAG GACGAAGCGG AGAGAGCCAA GACGGAGGAG GGAAGGGGC
2341 TGAGGACGAG ACCCCCATCC AGCCTCCAGG GCATTGCTCA GCCCGCTTGG CCACCCGGGG
2401 CCCTGCTTCC TCACACTGGC GGGTTTTTAC CCGGAACCGA GCCCATGGTG TTGGCTCTCA
2461 ACGTGACCCG CAGTCTGATC CCCTGTGAAG AGCCGGACAT CCCAGGCACA CGTGTGCGCC
2521 ACCTTCAGCA GGCATTCGGG TGCTGGGCTG GTGGCTCATC AGGCCTGGGC CCCACACTGA
2581 CAAGCGCCAG ATACGCCACA AATACCACTG TGTCAAATGC TTCAAGATA TATTTTGGG
2641 GAAACTATTT TTTAAACACT TTGGAATACA CTGGAAATCT TCAGGGAAAA ACACATTAA
2701 ACACTTTTTT TTTTAAGCCC

GENBANK ID: J02683

DEFINITION HUMAN ADP/ATP CARRIER PROTEIN MRNA, COMPLETE CDS.

VERSION J02683.1 GI:179246
CDS 70..966
/CODON_START=1

5 1 CCGCAGCGCC GTAGTCAAAC CGAACCCGGC CCAGTCCCGT CCTGCAGCAG TCTGCCTCCT
61 TCTTTCAACA TGACAGATGC CGCATTGTCC TTCGCCAAGG ACTTCCTGGC AGGTGGAGTG
121 GCCGCAGCCA TCTCCAAGAC GGCGGTAGCG CCCATCGAGC GGGTCAAGCT GCTGCTGCAG
181 GTGCAGCATG CCAGCAAGCA GATCACTGCA GATAAGCAAT ACAAAGGCAT TATAGACTGC
241 GTGGTCCGTA TTCCCAAGGA GCAGGAAGTT CTGTCCCTTCT GGC GCGGTAA CCTGGCCAAT
10 301 GTCATCAGAT ACTTCCCCAC CCAGGCTCTT AACTTCGCCT TCAAAGATAA ATACAAGCAG
361 ATCTTCCTGG GTGGTGTGGA CAAGAGAACC CAGTTTGGC GCTACTTTGC AGGGAATCTG
421 GCATCGGGTG GTGCCGAGG GGCCACATCC CTGTGTTTTG TGTACCCTCT TGATTTTGCC
481 CGTACCCGTC TAGCAGCTGA TGTGGGTAAA GCTGGAGCTG AAAGGGAATT CCGAGGCCTC
541 GGTGACTGCC TGGTTAAGAT CTACAAATCT GATGGGATTA AGGGCCTGTA CCAAGGCTTT
15 601 AACGTGTCTG TGCAGGGTAT TATCATCTAC CGAGCCGCCT ACTTCGGTAT CTATGACACT
661 GCAAAGGGAA TGCTTCCGGA TCCCAAGAAC ACTCACATCG TCATCAGCTG GATGATCGCA
721 CAGACTGTCA CTGCTGTTGC CGGGTTGACT TCCTATCCAT TTGACACCGT TCGCCGCCGC
781 ATGATGATGC AGTCAGGGCG CAAAGGAAC GACATCATGT ACACAGGCAC GCTTGACTGC
841 TGGCGGAAGA TTGCTCGTGA TGAAGGAGGC AAAGCTTTT TCAAGGGTGC ATGGTCCAAT
20 901 GTTCTCAGAG GCATGGGTGG TGCTTTTGTG CTTGTCTTGT ATGATGAAAT CAAGAAGTAC
961 ACATAAGTTA TTTCTTAGGA TTTTCCCCC TGTTGAACAG CATGTTGTAT TCTATAACAC
1021 AATCTTGAGC ATTCTTGACA GACTCCTGGC TGTGAGTTT TCAGTGGCAA CTACTTTACT
1081 GGTGAAAAT GGAAGCAAT AATATTCATC TGACCAGTTT TCTCTTAAAG CCATTTCCAT
1141 GCATGATGAT GATGGGACTC AATTGTATTT TTTATTTTCA TCACTCCTGA CTAAATAACA
25 1201 ATTTGGAGAA ATAAAAATAG TCTAAAAT

GENBANK ID: M22760.1

DEFINITION HOMO SAPIENS NUCLEAR-ENCODED MITOCHONDRIAL CYTOCHROME C OXIDASE VA
SUBUNIT MRNA, COMPLETE CDS.

30 VERSION M22760.1 GI:695359
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/CODON_START=1

35 1 GGGCGCCGCC ATCGCCGTCA TGCTGGGCGC CGCTCTCCGC CGCTGCGCTG TGGCCGCAAC
61 CACCCGGGCC GACCCTCGAG GCCTCCTGCA CTCCGCCCGG ACCCCCGGCC CCGCCGTGGC
121 TATCCAGTCA GTTCGCTGCT ATTCCCATGG GTCACAGGAG ACAGATGAGG AGTTTGATGC
181 TCGCTGGGTA ACATACTTCA ACAAGCCAGA TATAGATGCC TGGGAATTGC GTAAAGGGAT
241 AAACACACTT GTTACCTATG ATATGGTTCC AGAGCCCAA ATCATTGATG CTGCTTTGCG
301 GGCATGCAGA CGGTAAATG ATTTTGCTAG TCTAGTTTCA ATCCTAGAGG TTGTTAAGGA
40 361 CAAAGCAGGA CCTCATAAGG AAATCTACCC CTATGTCATC CAGGAACCTA GACCAACTTT
421 AAATGAACTG GGAATCTCCA CTCCGGAGGA ACTGGGCCTT GACAAAGTGT AAACCGCATG
481 GATGGGCTTC CCCAAGGATT TATTGACATT GCTACTTGAG TGTGAACAGT TACCTGGAAA
541 TACTGATGAT AACATATTAC CTTATTTTGA ACAAGTTTCC CTTTATTGAG TACCAAGCCA
45 601 TGTAATGGTA ACTTGACTT TAATAAAAGG GAAATGAGTT TGAAGT

GENBANK ID: M18079

DNA LINEAR

DEFINITION HUMAN, INTESTINAL FATTY ACID BINDING PROTEIN GENE, COMPLETE CDS, AND
AN ALU REPETITIVE ELEMENT.

50 VERSION M18079.1 GI:182351

55 1 GTAATATCTT GGGCAAGCCC TAGAGCTTCT TTCCTGACCC TTAGTTAATA AGATGTTATC
61 TGGTCACATT CAGTCACAAT AATAGACTCA TTTTAGTAAT AAACATCTTA AGACTAGTAA
121 TTAAACTCT TTAATTCACA CCAAGTTTCC TCCCAAGCT TGGCCTGTTT CTGGCTGGCA
181 GCCTGAAGTA GGGAAAGGAG AGATATGGTG ACCTTTTCTT TGTACCTTTC TAGCTACCCCT
241 CTATACCCCTG ACCCCACATA CATAATTGAG CTGTGGCTTC TGACTCTACT GGGTTTGGGG
301 ATGAGAGGCA GTGAGAGTAA AATGAAGGAG TGGTTTTAAT TAATGGCACA GCTAAACTG
361 GATTTTGTTC TCTCTGCACA TGGCAGATGT TTAAAGCTCA TTCTTTCTTT TATGCAAGTT
421 TTTACACCAT CCAGCCTCAT TTGTACCTCT TGAATTTTGT CTCAGTGGCC TATCACCATT
60 481 CAGGATCAAG AAAAAATCA ATGAGCACTT ATTGTGTGTC ATGCACCCCTA CAAAGTGCCA
541 GGATATTTAT CCAAACTCCT GGCAATGCTA AACACAATGC AAAAAGACAT ATTAGAAAAC
601 GAATCTTATT AACTTTAGCT TTTCAACTGT ATTTTCATCAT AAAGTCTTAC TTTACAAGAT
661 AATTGCTGTT GTGAAAAGG GAAAGGTCAT GGTCTCATTT CCCAGATGTT ATTTGATATA
721 TGCTATAAAT TATATTACCT CCAACATAGT CTGCACTTTG AACTTAGAAA AACAATCTTC
65 781 AGACGGCATG CATTCTAATT CTTGAAATAA GTATGCCAC AAAGTGTAGT TTAAGACAGA
841 ATAGGTATGC TTCTCATGTT TTAATTCAGT TGAATTTTCA AAGATCTCAG GAATGTACAG

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901 AACGAGAATT AAGAATTAAT AAGAATAAGA ATTAATTAAT TGCTTGACAT AGAGTAGTTA
 961 GGTGATTTCC TGAACTTTAA GCTTCCACAT CACAGTATGA AGTTGGTTCA AGATAAGAAA
 1021 TATAATAAAT TCTCGCCCAA GGACAGACCT GAATCTCTAG CTGCCTAGAG GCTGACTCAA
 1081 CTGAAATCAT GCGGTTTGAC AGCACTTGGA AGGTAGACCG GAGTGAAAAC TATGACAAGT
 1141 TCATGGAAAA AATGGGTAAA GACTTTATTT CTTTGTGGCT CATCTTTGC TTTCTTACAA
 1201 ACATTTTCTT TTCTAACTCC TAAATCTCTA GGAGATTACA GATAGCTTAC AGATAGCTCC
 1261 TGATGTGGTA GAGAGGGATC CAGAAGATGT TCAGAGGAGG GAAACCATAT TTTCCCTTCT
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 1681 ATAATTTGAT ACCAATACTC TGGCAGCCCA TATACTATAC AAGATAGGCA AACAAATTTG
 1741 TGTCATTCCC CTAAAAGAAA AATCTGCATC AATTATAGCT TACAGTTTAG GAACTCTAAG
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 1861 CTCATTTTGT TGTCAGAAAG AAATGCCACA AGAAGCAAAT AGAACTATAA AGTTCAAAAT
 1921 GTTAAAGCCA CTAAGAAAAA CAAAGGGGCA TTTAAGAAAA AAGAATACTG TATATGTGGA
 1981 ATTAAAGATG TGCTTCCTTA TAAATATATG AATATACATT TTAATCCTTC ATTTAATATT
 2041 TCTAGAATTT GATTACTTA ACATGAAAT GAACAGTTTG TTAATCTTAT TAAGGTTGCT
 2101 CAGCTCTAAG ATTCTATAAT TCTGTACTCT ACTTAATTTT TCTCAAGTTA TGGAAAAACA
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 3661 GTCAAATTTA TAGCTATTTT CAAAAGGCAA AAATTACTAC AAAACAATAA TTTTGTGCAC
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 3781 AAAATATTGT ATAGCTATTT TCTGATGCCT ATTTACTAAA GACAACCTAT ATATGTCAAA
 3841 TAATCAATGC CTATTTTAACT TGAAAATATA AATGACTACA AACCACATG TGTTTAAAAA
 3901 TGGCTGTATC CCATATCTGT ATAAATCTTG CTATCAAGTA CAAGAAAAAA TTGTATAAAC
 3961 TCATACTCAT ATAATATATA TGAATATATA ATATAAAAAAT AGTATAAACT CATATAGTAT
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4861 TATATGCTCA ACCAGAAAAC TTAGAAATAA GAAACACAAA TGTAATAATA GTATTTCCAT
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5041 ATGTACTACA TTAAATTAAA TTATTGTATT ACATTTTGT ACACATCAGT CATTCCCAGG
5101 CTTGGCTGAA AATCAGGATC ATCTGAGAAA CTAAACAAT TTCTGCATT TTAATCTCCA
5161 CTGTTATTCT ATTATATCAG AATCGCTAAT AGAACCAAGA ATTC

GENBANK ID: X16277

DNA LINEAR

10 DEFINITION HUMAN GENE FOR ORNITHINE DECARBOXYLASE ODC (EC 4.1.1.17).

VERSION X16277.1 GI:35137

MRNA JOIN(795..1001,3858..3967,4073..4191,4475..4648,

4855..5027,5286..5420,5551..5632,5809..5892,6948..7110,

7193..7305,7399..7613,8254..8740)

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1 GGATCCGGGT CCCCTCACGC TCCTGGCTGA GTCCCTGGCT TCACAGGGGA AACTACCTCC
61 GCAGGCCAGG ACCCATCTAG TTACAGGATA CCTCGATGTT ACAAAGACGA GGCTTCCAGC
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121 GCGGGGGCGT GGAGGCGGCT GCCAGCCCTG CCCGCAGCGT GCTGGCGACC CCCGGGACGC
181 CCCTTCCCTC CCGCGCCTCT GCTCCCTAGC TGGTGGGAGC AGAGCGCACC GGGATCACTT
241 CCAGGTCCCT TGCACCGGAG GAATGGGCGG CAGCAGGGTC CGGAGTCGGC CCGGCGGGGC
301 CCACGTGGCC AGCACATCGG TCCTCCGCTC GCGATTTCCC TTTTCCGCTC TCGGGCACGA
361 GGTACTGAAC GCCAGGTGGA AGCACAGCTG TGCAGCTACA GGCTCTGCCG TTCAGCTGCC
421 GCGGGCCGGG GCGGGGGCCT GCGGCGTCGT GCGCGTGCGC GGACCAAGTC CAGGCGGGCG
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481 AGACCGCCGC AGGGCGGGGC GGGGCGAGGC GGCCGCAGGG CGGGGAGGGC GGGGAGAGGC
541 GGCCGCAGGG CGGGGAGGGC GGGGCGCGAA GCCGGGGGCG GGGGCCACGC GTGGGGCAGG
601 CGGTGCTCGG CTCGGCTGAC GTCGGCCCGC CGGCGCCCCA CCAGCTCCGC GCGGGCCCCG
661 GTTGGCCACC GCCGGGCCCC CGCCCCCTCC CCGGCCGTGT CCCGGCCGGA ACCGATCGTG
721 GCTGGTTTGA GCTGGTGCCT CTCCATGGCG ACCCGCCGGT GCTATAAGTA GGGAGCGGCG
781 TGCCGTGGGG CTTTGTCACT CCCTCCTGTA GCCGCCGCCG CCGCCGCCCG CCGCCCCCTC
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841 GCCAGCAGCT CCGGCGCCAC CTCGGGCGCG CGTCTCCGGC GGGCGGGAGC CAGGCGCTGA
901 CGGGCGCGGC GGGGCGGGCC GAGCGCTCCT GCGGCTGCGA CTCAGGCTCC GGCGTCTGCG
961 CTTCCCCATG GGGCTGGCCT GCGGCGCCTG GGCGCTCTGA GGTGAGGGAC TCCCCGGCCG
1021 CGGAGGAAGG GAGGGAGCGA GGGGCGGAGC CCGGGCGGGC TCGGGGCCCC GGGCCCCGGG
1081 CACGTGTGCG CGCGCCCTCG CCGGCGCTCA GAGACACGTG GTCGCCGAGC GGGCCACGAC
35
1141 CTTGAGGCGC CGCTTCCCTC CCGCCCGGGG TTCTCCCGCG GCTGGATAAG GGTGATCCGG
1201 GCGCCTCGTT CTGCCCCCGT CTTACAGCT CCGGGCTGGA GGGGCGTAGG GGAGACCCAC
1261 CCGGAGACCC TGCGGCCCCG CGCGGCGCTC TTTCCCAACC CTTCGGCGGC CGCGCGCTGG
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1441 TTATTTTCGC TGTGTCTACA GAGCAGATGA CACCAATTTG GAAACCCGCG AGAGTGGGTA
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1681 AGTCGAAATA ATTGGTGAAA GTGTAGAAGG CAGAACCTCT CAGGAATGTC TGGGGAGGAC
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1741 AAAGATGTG TTGGCTGACT TTGTTTAAAC ATAAAATTGG GCAGACTTTA ATTGATTTGT
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1981 AAAGTATTGG CGTATTAAAA AGAAATCAAA ACTTTCCAAG TTTAGGCCTG AACATTGCCT
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2041 TAAAAATATT TAATAAGGCC TCAAATGACC CAGTCCGAGA CTGCATGAGC CTATTTATTA
2101 TTAAATTGTA AATATTCTTC ATATAAACAA AAATATATAA CCATGTCTGT AACAAAAATG
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2221 GTTGACAATG CCAAGCAGTC ACAATAGATA GAGCTTTAAA GCAAATTCTA TGCATGGGTT
2281 TGGATTTATG ACAGGCCCGT CACCCTGGGC CTGTCAATAGT ACCCATGCC AGAGCAAAT
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2341 GTGTCCCGGA ACCATTGCCT GGCTCTGTG CCCGTAGGCT GCTGGCACTG AAGTGGGTTG
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DEFINITION H.SAPIENS RING4 CDNA.
VERSION X57522.1 GI:36060
CDS 31..2457
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GENBANK ID: D00017.1

DEFINITION HOMO SAPIENS MRNA FOR LIPOCORTIN II, COMPLETE CDS.

VERSION D00017.1 GI:219909

CDS 50..1069

/CODON_START=1

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GENBANK ID: M10277.1

DNA LINEAR

DEFINITION HUMAN CYTOPLASMIC BETA-ACTIN GENE, COMPLETE CDS.

VERSION M10277.1 GI:177967

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2641 ACGCCAACAC AGTGCTGTCT GCGGGCACCA CCATGTACCC TGGCATTGCC GACAGGATGC
2701 AGAAGGAGAT CACTGCCCTG GCACCCAGCA CAATGAAGAT CAAGGTGGGT GTCTTTCTCTG
2761 CCTGAGCTGA CCTGGGCAGG TCAGCTGTGG GGTCTGTGG TGTGTGGGGA GCTGTCACAT
2821 CCAGGGTCTT CACTGCCTGT CCCCTTCCCT CCTCAGATCA TTGCTCCTCC TGAGCGCAAG
2881 TACTCCGTGT GGATCGGCGG CTCCATCCTG GCCTCGCTGT CCACCTTCCA GCAGATGTGG
2941 ATCAGCAAGC AGGAGTATGA CGAGTCCGGC CCCTCCATCG TCCACCGCAA ATGCTTCTAG
3001 GCGGACTATG ACTTAGTTGC GTTACACCCT TTCTTGACAA AACCTAACTT GCGCAGAAAA
3061 CAAGATGAGA TTGGCATGGC TTTATTTGTT TTTTGTGTT TGTGTTGGTT TTTTTTTTTT
3121 TTTTGGCTTG ACTCAGGATT TAAAACTGG AACGGTGAAG GTGACAGCAG TCGGTTGGAG
3181 CGAGCATCCC CCAAAGTTCA CAATGTGGCC GAGGACTTTG ATTGCATTGT TGTTTTTTTA
3241 ATAGTCATTC CAAATATGAG ATGCATTGTT ACAGGAAGTC CCTTGCCATC CTAAAAGCCA
3301 CCCCCTTCT CTCTAAGGAG AATGGCCCGG TCCTCTCCCA AGTCCACACA GGGGAGGTGA
3361 TAGCATTGCT TTCGTGTAAA TTATGTAATG CAAAATTTTT TTAATCTTCG CCTTAATACT
3421 TTTTATTTTT GTTTTATTTT GAATGATGAG CCTTCGTGCC CCCCCTTCCC CCTTTTGTCTC
3481 CCCCCTTCTG AGATGTATGA AGGCTTTTGG TCTCCCTGGG AGTGGGTGGA GGCAGCCAGG
3541 GCTTACCTGT AACTGACTT GAGACCAGTT GAATAAAAGT GCACACCTTA AAAATGAGGC
3601 CAAGTGTGAC TTTGTGGTGT GGCTGGGTTG GGGGCAGCAG AGGGTG

GENBANK ID: XM_042788.1

DEFINITION HOMO SAPIENS ALDOLASE B, FRUCTOSE-BISPHOSPHATE (ALDOB), MRNA.

VERSION XM_042788.1 GI:14738248

CDS 126..1220

/CODON_START=1

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1 AAAAATCATGA TGAGAAGTCT ATAAAAATTTG TGTGCTACCA AAGATCTGTC TTATTTGGCA
61 GCTGCTGCCCT CACCCACAGC TTTTGATATC TAGGAGGACT CTTCTCTCCC AAACCTACCTG
121 TCACCATGGC CCACCGATTT CCAGCCCTCA CCCAGGAGCA GAAGAAGGAG CTCTCAGAAA
181 TTGCCCAGAG CATTGTTGCC AATGGAAAGG GGATCCTGGC TGCAGATGAA TCTGTAGGTA
241 CCATGGGGAA CCGCCTGCAG AGGATCAAGG TGGAAGAAC TGAAGAGAAC CGCCGGCAGT

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301 TCCGAGAAAT CCTCTTCTCT GTGGACAGTT CCATCAACCA GAGCATCGGG GGTGTGATCC
361 TTTTCCACGA GACCCCTCTAC CAGAAGGACA GCCAGGGAAA GCTGTTTCTA AACATCCTCA
421 AGGAAAAGGG GATCGTGGTG GGAATCAAGT TAGACCAAGG AGGTGCTCCT CTTGCAGGAA
481 CAAACAAAGA AACCACCATT CAAGGGCTTG ATGGCCTCTC AGAGCGCTGT GCTCAGTACA
541 AGAAAGATGG TGTTGACTTT GGGAAAGTGGC GTGCTGTGCT GAGGATTGCC GACCASTGTC
601 CATCCAGCCT CGCTATCCAG GAAAACGCCA ACGCCCTGGC TCGCTACGCC AGCATCTGTC
661 AGCAGAATGG ACTGGTACCT ATTGTTGAAC CAGAGGTAAT TCCTGATGGA GACCATGACC
721 TGGAACACTG CCAGTATGTT ACTGAGAAGG TCCTGGCTGC TGTCTACAAG GCCCTGAATG
781 ACCATCATGT TTACCTGGAG GGCACCCTGC TAAAGCCCAA CATGGTGACT GCTGGACATG
841 CCTGCACCAA GAAGTATACT CCAGAACAAG TAGCTATGGC CACCGTAACA GCTCTCCACC
901 GTACTGTTCC TGCAGCTGTT CCTGGCATCT GCTTTTTGTC TGGTGGCATG AGTGAAGAGG
961 ATGCCACTCT CAACCTCAAT GCTATCAACC TTTGCCCTCT ACCAAAGCCC TGGAAACTAA
1021 GTTCTCTCTTA TGGACGGGCC CTGCAGGCCA GTGCACTGGC TGCCTGGGGT GGCAAGGCTG
1081 CAAACAAGGA GGCAACCCAG GAGGCTTTTA TGAAGCGGGC CATGGCTAAC TGCCAGGCGG
1141 CCAAAGGACA GTATGTTTAC ACGGGTTCTT CTGGGGCTGC TTCCACCCAG TCGCTCTTCA
1201 CAGCCTGCTA TACCTACTAG GGTCCAATGC CCGCCAGCCT AGCTCCAGTG CTTCTAGTAG
1261 GAGGGCTGAA AGGGAGCAAC TTTTCTCCA ATCCTGGAAA TTCGACACAA TTAGATTGA
1321 ACTGCTGGAA ATACAACACA TGTTAAATCT TAAGTACAAG GGGGAAAAAA TAAATCAGTT
1381 ATTGAAACAT AAAAATGAAT ACCAAGGACC TGATCAAATT TCACACAGCA GTTTCCTTGC
1441 AACACTTTCA GCTCCCATG CTCCAGAATA CCCACCCAAG AAAATAATAG GCTTTAAAC
1501 AATATCGGCT CCTCATCCAA AGAACAACCTG CTGATTGAAA CACCTCATTA GCTGAGTGTA
1561 GAGAAGTGCA TCTTATGAAA CAGTCTTAGC AGTGGTAGGT TGGGAAGGAG ATAGCTGCAA
1621 CCAAAAAGA AATAAATATT CTATAAACCT TC

GENBANK ID: NM_005317.2
DEFINITION HOMO SAPIENS GRANZYME M (LYMPHOCYTE MET-ASE 1) (GZMM), MRNA.
VERSION NM_005317.2 GI:7108347
CDS 46..819
/CODON_START=1

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121 ATCATCGGGG GCCGGGAGGT GATCCCCAC TCGCGCCCGT ACATGGCCTC ACTGCAGAGA
181 AATGGCTCCC ACCTGTGCGG GGGTGTCTCT GTGCACCCAA AGTGGGTGCT GACGGCTGCC
241 CACTGCCTGG CCCAGCGGAT GGCCAGCTG AGGCTGGTGC TGGGGCTCCA CACCCTGGAC
301 AGCCCCGGTC TCACCTTCCA CATCAAGGCA GCCATCCAGC ACCCTCGCTA CAAGCCCGTC
361 CCTGCCCTGG AGAACGACCT CGCGCTGCTT CAGCTGGACG GGAAAGTGAA GCCCAGCCGG
421 ACCATCCGGC CGTTGGCCCT GCCCAGTAAG CGCCAGGTGG TGGCAGCAGG GACTCGGTGC
481 AGCATGGCCG GCTGGGGGCT GACCCACCAG GCGGGCGGCC TGTCCCGGGT GCTGCGGGAG
541 CTGGACCTCC AAGTGCTGGA CACCCGATG TGTAACAACA GCCGCTTCTG GAACGGCAGC
601 CTCTCCCCCA GCATGGTCTG CCTGGCGGCC GACTCCAAGG ACCAGGCTCC CTGCAAGGGT
661 GACTCGGGCG GGCCCTGGT GTGTGGCAA GGCCGGGTGT TGGCCGGAGT CCTGTCCTTC
721 AGCTCCAGGG TCTGCACTGA CATCTTCAAG CCTCCCGTGG CCACCGCTGT GCGCCTTAC
781 GTGTCCTGGA TCAGGAAGGT CACCGGCCGA TCGGCCTGAT GCCCTGGGGT GATGGGGACC
841 CCCTCGCTGT CTCCACAGGA CCCTTCCCCT CCAGGGGTGC AGTGGGGTGG GTGAGGACGG
901 GTGGGAGGGA CAGGGAGGGA CCAATAAATC ATAATGAAGA AACGCTC

GENBANK ID: XM_003595.2
DEFINITION HOMO SAPIENS GLUTAMYL AMINOPEPTIDASE (AMINOPEPTIDASE A)
(ENPEP), MRNA.
VERSION XM_003595.2 GI:13647140
CDS 1401..2957
/CODON_START=1

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1 TCCAATTTAA AAAGGAAGTC TGCTGACGTT AGTTAGTTAA ATTTAACATC TTTTATGTG
61 TAACACTTGA CTTTGAAGC AAAAATGAAC TTTGCGGAGA GAGAGGGCTC TAAGAGATAC
121 TGCATTCAA CGAAACATGT GGCCATTCTC TGTGCGGTGG TGGTGGGTGT AGGATTAATA
181 GTGGGACTTG CCGTGGGCTT GACCAGATCG TGTGACTCCA GCGGGGACGG CCGGCCGGGC
241 ACTGCGCCAG CTCCTTCCCA CCTGCCTTCT TCCACGGCCA GCCCTCAGG TCCTCTGCC
301 CAGGACCAGG ACATCTGCCC GGCCAGTGAG GATGAGAGCG GACAGTGGA AACTTTTGA
361 CTGCCGGACT TCGTCAACCC AGTCCACTAC GACCTGCACG TGAAGCCCCT GTTGGAGGAG
421 GACACCTACA CGGGCACCGT GAGCATCTCC ATCAACCTGA GCGTCCAC CCGGTACCTG
481 TGGCTGCACC TCCGGGAGAC CAGGATCACC CCGCTCCCGG AGCTGAAGAG GCCCTCTGGG
541 GACCAGGTGC AAGTCCGGAG GTGTTTCGAG TACAAAAAGC AGGAGTACGT GGTGGTCGAG
601 GCGGAGGAAG AGCTTACCCC CAGCAGTGGA GATGGCCTGT ATCTCCTGAC CATGGAGTTC

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661 GCCGGCTGGC TGAACGGCTC CCTCGTGGGA TTTTATAGAA CCACCTACAC GGAGAACGGA
721 CAAGTCAAGA GCATAGTGGC CACCGATCAT GAACCAACAG ATGCCAGGAA ATCTTTTCCT
781 TGTTTTGATG AGCCCAACAA AAAGGCAACT TATACAATAT CTATCACCCA TCCCAAAGAA
841 TACGGAGCAC TTTCAAATAT GCCAGTGGCG AAAGAAGAGT CAGTGGATGA TAAATGGACT
901 CGAACAACTT TTGAGAAGTC TGTCCCATG AGCACGTACC TGGTGTGCTT TGCTGTACAT
961 CAATTTGACT CTGTAAAGAG AATATCAAAT AGTGGAAAAC CTCTTACAAT TTATGTCCAG
1021 CCAGAGCAAA AGCACACAGC CGAATATGCT GCAAACATAA CTAAAAGTGT GTTTGATTAT
1081 TTTGAAGAAT ACTTTGCTAT GAATTATTCT CTTCCTAAAT TAGATAAAAT CGCTATTCCA
1141 GATTTTGGCA CTGGTGCCAT GGAGAACTGG GGACTCATCA CGTACAGAGA AACGAACCTG
1201 CTTTATGACC CTAAGGAATC AGCCTCATCA AACCAACAGA GGGTGGCCAC TGTGGTTGCC
1261 CATGAACCTG TGCATCAGTG GTTTGGAAAT ATTTGTGACCA TGGACTGGTG GGAAGACTTG
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1381 GACTGGCAAA TGGTGACCAA ATGTTACTTG AAGATGTATT ACCTGTTCAA GAGGATGATT
1441 CTTTGATGTC TTCGCATCCA ATTATTGTGA CTGTGACAAC CCCTGATGAA ATAACATCTG
1501 TTTTGTATGG AATATCCTAT AGCAAGGGAT CTTCTATTTT GAGAATGCTT GAAGACTGGA
1561 TAAAACCAGA GAATTTTCAA AAAGGATGTC AGATGTACTT GGAAAAATAC CAATTCAGA
1621 ATGCAAAAC TTCTGATTTT TGGGCAGCAC TGGAGAGGC AAGTAGGCTA CCAGTGAAAG
1681 AAGTAATGGA CACCTGGACC AGACAGATGG GTTATCCTGT GCTTAACGTG AACGGTGTCA
1741 AGAACATCAC ACAGAAACGC TTTTGTGTTG ACCCAAGAGC TAACCCTTCT CAGCCCCCTT
1801 CAGATCTTGG TTATACATGG AATATCCCAG TTAATGGAC TGAAGATAAT ATAACAAGCA
1861 GTGTGTTATT TAATAGGTCA GAAAAGAAG GAATCACTTT GAACTCCTCT AATCCTAGTG
1921 GAAATGCTTT TCTCAAATA AACCCAGATC ATATTGGGTT TTATCGTGTA AATTATGAAG
1981 TAGCAACTTG GGAATCGATA GCTCAGCGC TCTCCTTGAA CCACAAGACA TTTTCTTCAG
2041 CAGATCGTGC AAGTCTTATT GATGATGCTT TTGCCTTGGC AAGAGCTCAA CTTCTAGATT
2101 ATAAGGTGGC TTTGAACCTG ACCAAGTATC TCAAAAGGGA AGAGAATTTT TTACCATGGC
2161 AGAGAGTAAT TTCAGCTGTA ACCTACATCA TTAGCATGTT TGAAGATGAT AAAGAGCTAT
2221 ATCCTATGAT TGAGGAATAC TTCCAAGGTC AAGTGAAGCC TATTGCAGAT TCTCTGGGAT
2281 GGAATGATGC TGGAGACCAT GTCACAAAGT TACTCCGTTT CTCCGTGTTA GGGTTTGCCT
2341 GCAAGATGGG AGACAGAGAA GCCTTGAACA ATGCTTCCTC GTTATTTGAG CAGTGGCTAA
2401 ATGGGACTGT AAGCCTTCCC GTAAATCTCA GGCTTCTGGT GTATCGGTAT GGGATGCAGA
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2521 CTCAAGAAAA AGAAAACTG CTGTATGGAT TAGCATCAGT GAAGAACGTT ACTCTTTTGT
2581 CAAGGTATTT GGATTTGCTC AAGGACACGA ACCTTATTAA AACTCAGGAT GTGTTTACAG
2641 TCATTCGATA TATCTCATAT AACAGCTATG GGAAGAACAT GGCTGGAAT TGGATACAAC
2701 TCAACTGGGA CTATCTAGTC AACAGATATA CACTCAATAA CAGAAACCTT GGCCGAATTG
2761 TCACAATAGC AGAGCCATTC AACACTGAAC TGCAACTGTG GCAGATGGAG AGCTTTTTTG
2821 CAAAATATCC ACAAGCTGGA GCAGGAGAAA AACCTAGGGA ACAAGTGCTG GAAACAGTGA
2881 AAAACAATAT AGAGTGGCTA AAACAACATA GAAACACCAT CAGAGAATGG TTTTAAATT
2941 TACTTGAGAG TGGTTAATGT ATTCAAATGT TAGAGTTTAA TTTTGTGAAT CTATTGTTTC

GENBANK ID: U34683.1

DEFINITION HUMAN GLUTATHIONE SYNTHETASE MRNA, COMPLETE CDS.

VERSION U34683.1 GI:1236349

CDS 41..1465

/CODON_START=1

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1 GGGAGAACCG TTCGCGGAGG AAAGGCGAAC TAGTGTGGG ATGGCCACCA ACTGGGGGAG
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121 GGCTGAGGGA GTATTGCTGA GGACCTCACA GGAGCCCACT TCCTCGGAGG TGGTGAGCTA
181 TGCCCCATTC ACGCTCTTCC CCTCACTGGT CCCAGTGCC CTGCTGGAGC AAGCCTATGC
241 TGTGCAGATG GACTTCAACC TGCTAGTGGG TGCTGTCAGC CAGAACGCTG CCTTCCTGGA
301 GCAAACTCTT TCCAGCACCA TCAAACAGGA TGAATTTACC GCTCGTCTCT TTGACATCCA
361 CAAGCAAGTC CTAAAAGAGG GCATTGCCCA GACTGTGTTC CTGGGCCTGA ATCGCTCAGA
421 CTACATGTTT CAGCGCAGCG CAGATGGCTC CCCAGCCCTG AAACAGATCG AAATCAACAC
481 CATCTCTGCC AGCTTTGGGG GCCTGGCCTC CCGGACCCCA GCTGTGCACC GACATGTTCT
541 CAGTGTCTTG AGTAAGACCA AAGAAGCTGG CAAGATCCTC TCTAATAATC CCAGCAAGGG
601 ACTGGCCCTG GGAATTGCCA AAGCCTGGGA GCTCTACGGC TCACCCAATG CTCTGGTGCT
661 ACTGATTGCT CAAGAGAAGG AAAGAAACAT ATTTGACCAG CGTGCCATAG AGAATGAGCT
721 ACTGGCCAGG AACATCCATG TGATCCGACG AACATTTGAA GATATCTCTG AAAAGGGGCT
781 TCTGGACCAA GACCGAAGGC TGTTTGTGGA TGGCCAGGAA ATTGCTGTGG TTTACTTCCG
841 GGATGGCTAC ATGCCTCGTC AGTACAGTCT ACAGAATTGG GAAGCACGTC TACTGCTGGA
901 GAGGTCACAT GCTGCCAAGT GCCCAGACAT TGCCACCCAG CTGGCTGGGA CTAAGAAGGT
961 GCAGCAGGAG CTAAGCAGGC CGGGCATGCT GGAGATGTTG CTCCCTGGCC AGCCTGAGGC
1021 TGTGGCCCCG CTCCGCGCCA CCTTTGCTGG CCTTACTCA CTGGATGTGG GTGAAGAAGG
1081 GGACCAGGCC ATCGCCGAGG CCCTTGCTGC CCTAGCCGG TTTGTGCTAA AGCCCCAGAG
1141 AGAGGGTGGA GGTAACAACC TATATGGGGA GGAAATGGTA CAGGCCCTGA AACAGCTGAA

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1201 GGACAGTGAG GAGAGGGCCT CCTACATCCT CATGGAGAAG ATCGAACCTG AGCCTTTTGA
1261 GAATTGCCTG CTACGGCCTG GCAGCCCTGC CCGAGTGGTC CAGTGCATTT CAGAGCTGGG
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1381 TCTACTTCGA ACCAAAGCCA TCGAGCATGC AGATGGTGGT GTGGCAGCGG GAGTGGCAGT
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1501 TATTTGTCAT TCCTCTCCTA GCCCTCCTGA GGGGTATCCT CCTAAAGACC TCCAAAGTTT
1561 TTATGGAAGG GTAAATACTG GTACCTTCCC CCAGCTTTCC ATCTGAGGAC CAGAAAAGTT
1621 GTGTCTCCCT TAGATGAGAT CTAGACGCCC CCAAATCCTT GAGATGTGGG TATAGCTCAG
1681 GGTAAGCTGC TCTGAGGTAA AGGTCCATGA ACCCTGCCCC ACTCCTGTCA GCCCCTCATC
1741 AGCCTTTTCA GCAGGTTCCA GTGCCTGACT TGGGATAGGA CTGAGTGGTA GGAGGAGGGG
1801 GAGTGGAGGG GCATAGCCTT TCCCTAATTC TGCCTTAAAT AAAACTGCAT TGCTGT

GENBANK ID: AF035429.1

DNA LINEAR

15
DEFINITION HOMO SAPIENS CYTOCHROME OXIDASE SUBUNIT I (COI) AND SUBUNIT II
(COII) PSEUDOGENES, COMPLETE SEQUENCE.
VERSION AF035429.1 GI:2665724

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1 AATATGAAAA TCACCTCGGA GCTGGTAAAA AGAGGCTTAA CCCCTGTCTT TAGATTTACA
61 GTCCAATGCT TCACTCAGCC ATTTTACCTC ACCCCCACTG ATGTTGCGCG ACCGTTGACT
121 ATTCTCTACA AACCACAAAG ACATTGGAAC ACTATACCTA TTATTCGGCG CATGAGCTGG
181 AGTCCTAGGC ACAGCTCTAA GCCTCCTTAT TCGAGCCGAA CTGGGCCAGC CAGGCAACCT
241 TCTAGGTAAC GACCACATCT ACAACGTTAT CGTCACAGCC CATGCATTTG TAATAATCTT
301 CTTCATAGTA ATACCCATCA TAATCGGAGG CTTTGGCAAC TGACTAGTTC CCCTAATAAT
361 CGGTGCCCCC GATATGGCGT TTCCCCGCAT AAACAACATA AGCTTCTGAC TCTTACCCCC
421 CTCTCTCCTA CTCCTGCTTG CATCTGCTAT AGTGGAGGCC GGCGCAGGAA CAGGTTGAAC
481 AGTCTACCCT CCCTTGGCAG GGAACACTC CCACCCTGGA GCCTCCGTAG ACCTAACCAT
541 CTTCTCCTTA CACCTAGCAG GTATCTCCTC TATCTTAGGA GCCATCAATT TCATCACAAC
601 AATTATTAAT ATAAAACCCC CTGCCATAAC CCAATACCAA ACGCCCCCTT TCGTCTGATC
661 CGTCCTAATC ACAGCAGTCT TACTTCTCCT ATCTCTCCCA GTCCTAGCCG CTGGCATCAC
721 TATACTACTA ACAGACCGTA ACCTCAACAC CACCTTCTTC GACCCAGCCG GAGGAGGAGA
781 CCCCATCTA TACCAACACC TATTCTGATT TTTCGGTCAC CCTGAAGTTT ATATTCTCAT
841 CCTACCAGGC TTCGGAATAA TCTCCCATAT TGTAACCTAC TACTCCGGA AAAAAGAACC
901 ATTTGGATAC ATAGGTATGG TCTGAGCTAT GATATCAATT GGCTTCCTAG GGTTCATCGT
961 GTGAGCACAC CATATATTTA CAGTAGGAAT AGACGTAGAC ACACGAGCAT ATTTACCTC
1021 CGCTACCATA ATCATCGCTA TCCCCACCGG CGTCAAAGTA TTTAGCTGAC TCGCCACACT
1081 CCACGGAAGC AATATGAAAT GATCTGCTGC AGTGCTCTGA GCCCTAGGAT TTATTTTCT
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1321 AACCTACGCC AAAATCCATT TCGCTATCAT ATTCATCGGC GTAAATCTAA CTTTCTTCCC
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1501 AATAATTTTC ATAATTTGAG AAGCCTTCGC TTCGAAGCGA AAAGTCCTAA TAGTAGAAGA
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1621 ACCCGTATAC ATAAAATCTA GACAAAAAAG GAAGGAATCG AACCCCCAA AGCTGGTTTC
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1861 GCCCTCATA TCATTTTCTT TATCTGCTTC CTAGTCCTGT ACGCCCTTTT CCTAACACTC
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1981 ATCCTGCCCC CCATCATCCT AGTCCTTATC GCCCTCCCAT CCCTACGCAT CCTTTACATA
2041 ACAGACGAGG TCAACGATCC CTCCTTTACC ATCAAATCAA TTGGCCATCA ATGGTACTGA
2101 ACCTACGAAT ACACCGACTA CGGCGGACTA ATCTTCAACT CCTACATACT TCCCCATTA
2161 TTCTTAGAAC CAGGCGACCT GCGACTCCTT GACGTTGACA ATCGAGTAGT ACTCCCGGTT
2221 GAAGCCCCCA TTCGTATAAT AATTACATCA CAAGACGCTT TACACTCATG AGCTGTCCCC
2281 ACATTAGGCT TAAAAACAGA TGCAATTCCC GGACGTCTAA ACCAAACCAC TTTCACTGCT
2341 ACACGACCAG GGGTATACTA CGGCCAATGC TCTGAAATCT GTGGAGCAAA CCAGTTTTAT
2401 GCCCATCGTC CTAGAATTAA TTCCCCTAAA AATCTTTGAA ATAGGGCCTG TATTTACCCT
2461 ATAGCACCCC CTCTACCCCC TCTAGAGCCC ACTGTAAAGC TAACTTAGCA TTAACCTTTT
2521 AAGTTAAAGA TTAAGAGAAC CAACACCTCT TTACAGTGAA ATGCCCCAAC TAAATACTA

GENBANK ID: NM_024409.1

65
DEFINITION HOMO SAPIENS NATRIURETIC PEPTIDE PRECURSOR C (NPPC), MRNA.
VERSION NM_024409.1 GI:13249345

CDS 1..381
/CODON_START=1

5 1 ATGCATCTCT CCCAGCTGCT GGCCTGCGCC CTGCTGCTCA CGCTGCTCTC CCTCCGGCCC
61 TCCGAAGCCA AGCCCGGGGC GCCGCCGAAG GTCCCGCGAA CCCC GCCGGC AGAGGAGCTG
121 GCCGAGCCGC AGGCTGCGGG CGGCGGTCAG AAGAAGGGCG ACAAGGCTCC CGGGGGCGGG
181 GGCGCCAATC TCAAGGGCGA CCGGTGCGCA CTGCTCCGGG ACCTGCGCGT GGACACCAAG
241 TCGCGGGCAG CGTGGGCTCG CTTTCTGCAA GAGCACCCCA ACGCGCGCAA ATACAAAGGA
301 GCCAACAAGA AGGGCTTGTG CAAGGGCTGC TTCGGCCTCA AGCTGGACCG AATCGGCTCC
10 361 ATGAGCGGCC TGGGATGTTA G

GENBANK ID: M37763.1
DNA LINEAR
DEFINITION HUMAN NEUROTROPHIN-3 (NT-3) GENE, COMPLETE CDS.
15 VERSION M37763.1 GI:189300
CDS 76..849
/CODON_START=1
GENE 76..849
20 MAT PEPTIDE 130..846

1 TAACACAGAC TCAGCTGCCA GAGCCTGCTC TTAACACCTG TGTTTCCTTT TCAGATCTTA
61 CAGGTGAACA AGGTGATGTC CATCTTGTTT TATGTGATAT TTCTCGCTTA TCTCCGTGGC
121 ATCCAAGGTA ACAACATGGA TCAAAGGAGT TTGCCAGAAG ACTCGCTCAA TTCCCTCATT
181 ATTAAGCTGA TCCAGGCAGA TATTTTGAAA AACAAGCTCT CCAAGCAGAT GGTGGACGTT
25 241 AAGGAAAATT ACCAGAGCAC CCTGCCCAA GCTGAGGCTC CCCGAGAGCC GGAGCGGGGA
301 GGGCCCGCCA AGTCAGCATT CCAGCCGGTG ATTGCAATGG ACACCGAACT GCTGCGACAA
361 CAGAGACGCT ACAACTCACC GCGGGTCCTG CTGAGCGACA GCACCCCTT GGAGCCCCCG
421 CCCTTGATAT TCATGGAGGA TTACGTGGGC AGCCCCGTGG TGGCGAACAG AACATCACGG
481 CGGAAACGGT ACGCGGAGCA TAAGAGTCAC CGAGGGGAGT ACTCGGTATG TGACAGTGAG
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601 CTGGGGGAGA TCAAAACGGG CAACTCTCCC GTCAAACAAT ATTTTATGA AACCGGATGT
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721 CAGTGCAAAA CATCCCAAAC CTACGTCCGA GCACTGACTT CAGAGAACAA TAAACTCGTG
781 GGCTGGCGGT GGATACGGAT AGACACGTCC TGTGTGTGTG CCTTGTGCGAG AAAAATCGGA
35 841 AGAACATGAA TTGGCATCTC TCCCCATATA TAAATTATTA CTTTAAATTA TATGATATGC
901 ATGTAGCATA TAAATGTTTA TATTGTTTTT ATATATTATA AGTTGACCTT TATTTATTAA
961 ACTTCAGCAA CCCTACAGTA TATAAGCTTT TTTCTCAATA AAATCAGTGT GCTTGCCCTC

GENBANK ID: NM_000932.1
40 DEFINITION HOMO SAPIENS PHOSPHOLIPASE C, BETA 3
(PHOSPHATIDYLINOSITOL-SPECIFIC) (PLCB3), MRNA.
VERSION NM_000932.1 GI:11386138
CDS 1..3705

45 1 ATGGCGGGCG CCCAGCCCGG CGTCCACGCG CTGCAGTTGG AGCCGCCAC CGTGGTGGAG
61 ACCCTGCGGC GCGGGAGTAA GTTCATCAAA TGGGACGAGG AGACCTCCAG TCGGAACCTG
121 GTGACCCTGC GTGTGGACCC CAATGGCTTC TTCTTGTAAT GGACGGGCCC CAACATGGAG
181 GTGGACACAC TGGACATCAG TTCCATCAGG GACACACGGA CAGGCCGGTA CGCCCGCCTG
241 CCCAAGGACC CCAAGATCCG GGAAGTTCTG GGCTTTGGGG GTCCCGATGC CCGGCTGGAG
50 301 GAGAAGCTGA TGACGGTGGT GTCTGGGCCA GACCCGGTGA ACACAGTGTT CTTGAACTTC
361 ATGGCCGTGC AGGATGACAC AGCCAAGGTC TGGTCTGAGG AGCTATTCAA GCTGGCTATG
421 AACATCCTGG CTCAGAACGC CTCCCGGAAC ACCTTCCTGC GCAAAGCATA CACGAAGCTG
481 AAGCTGCAGG TGAACCAGGA TGGTCGGATC CCCGTCAAGA ACATCCTGAA GATGTTCTCA
541 GCAGACAAGA AGCGGTGGA GACTGCGCTG GAATCCTGTG GCCTCAAATT CAACCGGAGT
55 601 GAGTCCATCC GGCTPGATGA GTTTTCCTTG GAAATCTTTG AGCGGTTTCT GAACAAGCTG
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781 GAAGTGCTGT ACCCGCCCCT GCGGCCCTCC CAGGCCCGGC TGCTCATCGA AAAGTATGAG
841 CCCAACCAGC AGTTTCTGGA GCGAGACCTG ATGTCCATGG AGGGCTTTAG CCGCTACCTG
60 901 GGAGGCGAGG AGAATGGCAT CCTGCCCTG GAAGCCCTGG ATCTGAGCAC GGACATGACC
961 CAGCCACTGA GTGCTACTT CATCAACTCC TCGCATAACA CCTATCTCAC TGCGGGGCAG
1021 CTGGCTGGGA CCTCGTCGGT GGAGATGTAC CGCCAGGCAC TACTATGGGG CTGCCGCTGC
1081 GTGGAGCTGG ACGTGTGGAA GGGACGGCCG CCTGAGGAGG AACCCTTCAT TACCCACGGC
1141 TTCACCATGA CCACAGAGGT GCCTCTGCGC GACGTGCTGG AGGCCATTGC CGAGACTGCC
65 1201 TTCAAGACCT CGCCCTACCC CGTCATCCTC TCCTTCGAGA ACCATGTGGA CTCGGCAAAG
1261 CAACAGGCAA AGATGGCTGA GTACTGCCGC TCCATCTTTG GAGACGCGCT ACTCATCGAG

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1321 CCTCTGGACA AGTACCCGCT GGCCCCAGGC GTTCCCCTGC CCAGCCCCCA GGACCTGATG
1381 GGCCGTATCC TGGTGAAGAA CAAGAAGCGG CACCGACCCA GCGCAGGTGG CCCAGACAGC
1441 GCCGGGCGCA AGCGGCCCTT GGAGCAGAGC AATTCTGCCC TGAGCGAGAG CTCCGCGGCC
1501 ACCGAGCCCT CCTCCCCGCA GCTGGGGTCT CCCAGCTCTG ACAGCTGCCC AGGCCTGAGC
1561 AATGGGGAGG AGGTAGGGCT TGAGAAGCCC AGCCTGGAGC CTCAGAAGTC TCTGGGTGAC
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1681 GATGAGGAAG AGGAGGAACA GACAGACCCC AAAAAGCCAA CTACAGATGA GGGCACAGCC
1741 AGCAGCGAGG TGAATGCCAC TGAGGAGATG TCCACGCTTG TCAACTACAT CGAACCTGTC
1801 AAGTTCAAGT CCTTTGAGGC TGCTCGAAAG AGGAACAAAT GCTTCGAGAT GTCGTCCTTT
1861 GTGGAGACCA AGGCCATGGA GCAACTGACC AAGAGCCCCA TGGAGTTTGT GGAATACAAC
1921 AAGCAGCAGC TCAGCCGCAT CTACCCCAAG GGCACCCGCG TGGACTCCTC CAACTACATG
1981 CCCCAGCTCT TCTGGAACGT AGGGTGCCAG CTTGTTGCGC TCAACTTCCA GACCCTCGAT
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2101 AAGCCGGAGT TCATGCGGCG GCCGGACAAG TCCTTCGACC CCTTCACTGA GGTCACTGTC
2161 GATGGCATCG TGGCCAATGC CTTGCGGGTC AAGGTGATCT CAGGGCAGTT CCTGTCCGAC
2221 AGGAAGGTGG GCATCTACGT GGAGGTGGAC ATGTTTGGCC TCCCTGTTGA TACGCGGCGC
2281 AAGTACCGCA CCCGGACCTC TCAGGGGAAC TCGTTCAACC CCGTGTGGGA CGAAGAGCCC
2341 TTCGACTTCC CCAAGGTGGT GCTGCCACG CTGGCTTCAC TTCGCATTGC AGCCTTTGAG
2401 GAGGGGGGTA AATTCGTAGG GCACCGGATC CTGCCTGTCT CTGCCATCCG CTCCGGATAC
2461 CACTACGTCT GCCTGCGGAA CGAGGCCAAC CAACCGCTGT GCCTGCCGGC CCTGCTCATC
2521 TACACCGAAG CCTCGGACTA CATTCCAGAC GACCACCAGG ACTATGCGGA GGCCCTGATC
2581 AACCCCATTA AGCAGCTCAG CTGATGGAC CAGAGGGCCC GGCAGCTGGC CGCCCTCATT
2641 GGGGAGAGTG AGGCTCAGGC TGGCCAAGAG ACGTGCCAGG ACACCCAGTC TCAGCAGCTG
2701 GGGTCTCAGC CGTCTCAAA CCCCACCCCC AGCCCACTGG ATGCCTCCCC CCGCCGGCCC
2761 CCTGGCCCCA CCACCTCCCC TGCCAGCACC TCCCTCAGCA GCCCAGGGCA GCGTGATGAT
2821 CTCATCGCCA GCATCCTCTC AGAGGTGGCC CCCACCCGCG TGGATGAGCT CCGAGGTCAC
2881 AAGGCTCTGG TCAAGCTCCG GAGCCGGCAA GAGCGAGACC TGCGGGAGCT GCGCAAGAAG
2941 CATCAGCGGA AGGCAGTCAC CCTCACCCGC CGCCTGCTGG ATGGCCTGGC TCAGGCACAG
3001 GCTGAGGGCA GGTGCCGGCT GCGGCCAGGT GCCCTAGGTG GGGCCGCTGA TGTGGAGGAC
3061 ACGAAGGAGG GGGAGGACGA GGCAAAGCGG TATCAGGAGT TCCAGAACAG ACAGGTGCAG
3121 AGCCTGCTGG AGCTGCGGGA GGCCCAGGTG GACGCAGAGG CCCAGCGGAG GCTGGAACAC
3181 CTGAGACAGG CTCTGCAGCG GCTCAGGGAG GTCGCTCTTG ATGCAAACAC AACTCAGTTC
3241 AAGAGGCTGA AAGAGATGAA CGAGAGGGAG AAGAAGGAGC TGCAGAAGAT CCTGGACAGA
3301 AAGCGCCATA ACAGCATCTC GGAGGCCAAG ATGAGGGACA AGCATAAGAA GGAGGCGGAA
3361 CTGACGGAGA TTAACCGTCG GCACATCACT GAGTCAGTCA ACTCCATCCG TCGGCTGGAG
3421 GAGGCCCAGA AGCAGCGGCA TGACCGTCTT GTGGCTGGGC AGCAGCAGGT CCTGCAACAG
3481 CTGGCAGAAG AGGAGCCCAA GCTGCTGGCC CAGCTGGCCC AGGAGTGTCG GGAGCAGCGG
3541 GCGAGGCTCC CCCAGGAGAT CCGCCGGAGC CTGCTGGGCG AGATGCCGGA GGGGCTGGGG
3601 GACGGGCCCTC TGGTGGCCTG TGCCAGCAAC GGTACGCAC CCGGGAGCAG CGGGCACCTG
3661 TCGGGCGCTG ACTCGGAGAG CCAGGAGGAG AACACGCAGC TCTGA

GENBANK ID: D13119.1

DEFINITION HOMO SAPIENS P2 MRNA FOR ATP SYNTHASE SUBUNIT C, COMPLETE CDS.

VERSION D13119.1 GI:285909

CDS 31..456

/CODON_START=1

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1 TCTCCTGCCA CAGCTCCTCA CCCCCTGAAA ATGTTGCGCT GCTCCAAGTT TGTCTCCACT
61 CCCTCCTTGG TCAAGAGCAC CTCACAGCTG CTGAGCCGTC CGCTATCTGC AGTGGTGCTG
121 AAACGACCGG AGATACTGAC AGATGAGAGC CTCAGCAGCT TGGCAGTCTC ATGTCCCCTT
181 ACCTCACTTG TCTCTAGCCG CAGCTTCCAA ACCAGCGCCA TTCAAGGGA CATCGACACA
241 GCAGCCAAGT TCATTGGAGC TGGGGCTGCC ACAGTTGGGG TGGCTGGTTC TGGGGCTGGG
301 ATTGGAAGTG TGTTTGGGAG CCTCATCATT GGTATGCCA GGAACCCTTC TCTGAAGCAA
361 CAGCTCTTCT CCTACGCCAT TCTGGGCTTT GCCCTCTCGG AGGCCATGGG GCTCTTTTGT
421 CTGATGGTAG CTTTTCTCAT CCTCTTTGCC ATGTGAAGGA GCCGTCTCCA CCTCCCATAG
481 TTCTCCGCG TCTGGTTGGC CCGGTGTGTT CCTTTTCTA TACCTCCCCA GGCAGCCTGG
541 GGAACGTGGT TGGCTCAGGG TTTGACAGAG AAAAGACAAA TAAATACTGT ATTAATAAG

GENBANK ID: NM_004530.1

DEFINITION HOMO SAPIENS MATRIX METALLOPROTEINASE 2 (GELATINASE A, 72KD GELATINASE, 72KD TYPE IV COLLAGENASE) (MMP2), MRNA.

VERSION NM_004530.1 GI:11342665

CDS 290..2272

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121 CGGGGGCCCG ACCATGAGCC GCTGAGCCGG GCAAACCCCA GGCCACCGAG CCAGCGGACC
181 CTCGGAGCGC AGCCCTGCGC CGCGGACCAG GCTCCAACCA GGCGGCGAGG CGGCCACACG
241 CACCGAGCCA GCGACCCCGG GCGGACGCGC GGGGCCAGGG AGCGCTACGA TGGAGGCGCT
301 AATGGCCCGG GCGCGCTCA CGGGTCCCCT CATCAAGTTC CCCGGCGATG TCGCCCCCAA
361 GAGCCACGCC GCCGCCGCGC CGTCGCCCAT CATCAAGTTC CCCGGCGATG TCGCCCCCAA
421 AACGGACAAA GAGTTGGCAG TGCAATACCT GAACACCTTC TATGGCTGCC CCAAGGAGAG
481 CTGCAACCTG TTTGTGCTGA AGGACACACT AAAGAAGATG CAGAAGTTCT TTGGACTGCC
541 CCAGACAGGT GATCTTGACC AGAATACCAT CGAGACCATG CGGAAGCCAC GCTGCGGCAA
601 CCCAGATGTG GCCAACTACA ACTTCTTCCC TCGCAAGCCC AAGTGGGACA AGAACCAGAT
661 CACATACAGG ATCATTGGCT ACACACCTGA TCTGGACCCA GAGACAGTGG ATGATGCCTT
721 TGCTCGTGCC TTCCAAGTCT GGAGCGATGT GACCCCACTG CGGTFTTCTC GAATCCATGA
781 TGGAGAGGCA GACATCATGA TCAACTTTGG CCGCTGGGAG CATGGCGATG GATACCCCTT
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901 CTCCCATTTT GATGACGATG AGCTATGGAC CTTGGGAGAA GGCCAAGTGG TCCGTGTGAA
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1021 CAACAGCTGC ACTGATACTG GCCGACGCGA TGGCTTCCTC TGGTGCTCCA CCACCTACAA
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1201 CTGCACCACT GAGGGCCGCA CGGATGGCTA CCGCTGGTGC GGCACCACTG AGGACTACGA
1261 CCGCGACAAG AAGTATGGCT TCTGCCCTGA GACCGCCATG TCCACTGTTG GTGGGAACCTC
1321 AGAAGGTGCC CCTGTGTCT TCCCCTTCAC TTTCTGGGC AACAAATATG AGAGCTGCAC
1381 CAGCGCCGGC CGCAGTGACG GAAAGATGTG GTGTGCGACC ACAGCCAACCT ACGATGACGA
1441 CCGCAAGTGG GGCTTCTGCC CTGACCAAGG GTACAGCCTG TTCCTCGTGG CAGCCCACGA
1501 GTTTGGCCAC GCCATGGGGC TGGAGCACTC CCAAGACCCT GGGGCCCTGA TGGCACCCAT
1561 TTACACCTAC ACCAAGAAGT TCCGTCTGTC CCAGGATGAC ATCAAGGGCA TTCAGGAGCT
1621 CTATGGGGCC TCTCCTGACA TTGACCTTGG CACCGGCCCC ACCCCACAC TGGGCCCTGT
1681 CACTCCTGAG ATCTGCAAAAC AGGACATTGT ATTTGATGGC ATCGCTCAGA TCCGTGGTGA
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1861 CGAGGCCCCA CAGGAGGAGA AGGCTGTGTT CTTTGCAGGG AATGAATACT GGATCTACTC
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2101 GCTCATCGCA GATGCCTGGA ATGCCATCCC CGATAACCTG GATGCCGTCG TGGACCTGCA
2161 GGGCGGCGGT CACAGCTACT TCTTCAAGGG TGCCATTATC CTGAAGCTGG AGAACCAGAG
2221 TCTGAAGAGC GTGAAGTTTG GAAGCATCAA ATCCGACTGG CTAGGCTGCT GAGCTGGCCC
2281 TGGCTCCAC AGGCCCTTCC TCTCCACTGC CTTGATACA CCGGGCCTGG AGAAGTAGAG
2341 AAGGACCCGG AGGGGCTTGG CAGCCGTGCC TTCAGCTCTA CAGCTAATCA GCATTCTCAC
2401 TCCTACCTGG TAATTTAAGA TTCCAGAGAG TGGCTCCTCC CGGTGCCCAA GAATAGATGC
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2521 CCTAAAGAGA TCCTTTGATA TTTTCAACGC AGCCCTGCTT TGGGCTGCCC TGGTGTGCTC
2581 AACTTTCAGG CTCTTCTCCT TTCACAACCT TCTGTGGCTC ACAGAACCCT TGGAGCCAAT
2641 GGAGACTGTC TCAAGAGGGC ACTGGTGGCC CGACAGCCTG GCACAGGGCA GTGGGACAGG
2701 GCATGGCCAG GTGGCCACTC CAGACCCCTG GCTTTTCACT GCTGGCTGCC TTAGAACCCT
2761 TCTTACATTA GCAGTTTGCT TTGTATGCAC TTTGTTTTTT TCTTTGGGTC TTGTTTTTTT
2821 TTTCCACTTA GAAATTGCAT TTCCTGACAG AAGGACTCAG GTTGTCTGAA GTCAGTGCAC
2881 AGTGCATCTC AGCCACATA GTGATGGTTC CCCTGTTTAC TCTACTTAGC ATGTCCCTAC
2941 CGAGTCTCTT CTCCACTGGA TGGAGGAAAA CCAAGCCGTG GCTTCCCGCT CAGCCCTCCC
3001 TGCCCCCTCCC TTCAACCATT CCCCATGGGA AATGTCAACA AGTATGAATA AAGACACCTA
3061 CTGAGTGGC

GENBANK ID: NM 000852.2

DEFINITION HOMO SAPIENS GLUTATHIONE S-TRANSFERASE PI (GSTP1), MRNA.

VERSION NM_000852.2 GI:6552334

CDS 30..662

1 GGAGTTTCGC CGCCGCAGTC TTCGCCACCA TGCCGCCCTA CACCGTGGTC TATTTCCCAG
61 TTCGAGGCCG CTGCGCGGCC CTGCGCATGC TGCTGGCAGA TCAGGGCCAG AGCTGGAAGG
121 AGGAGGTGGT GACCGTGGAG ACGTGGCAGG AGGGCTCACT CAAAGCCTCC TGCTTATACG
181 GGCAGTCCC CAAGTTCCAG GACGGAGACC TCACCCTGTA CCAGTCCAAT ACCATCCTGC
241 GTCACCTGGG CCGCACCTTT GGGCTCTATG GGAAGGACCA GCAGGAGGCA GCCCTGGTGG
301 ACATGGTGAA TGACGGCGTG GAGGACCTCC GCTGCAAATA CATCTCCCTC ATCTACACCA
361 ACTATGAGGC GGGCAAGGAT GACTATGTGA AGGCACTGCC CGGGCAACTG AAGCCTTTTG
421 AGACCCTGCT GTCCAGAAC CAGGGAGGCA AGACCTTCAT TGTGGGAGAC CAGATCTCCT

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481 TCGCTGACTA CAACCTGCTG GACTTGCTGC TGATCCATGA GGTCTTAGCC CCTGGCTGCC
541 TGGATGCGTT CCCCTGCTC TCAGCATATG TGGGGCGCCT CAGCGCCCGG CCCAAGCTCA
601 AGGCCTTCCT GGCCTCCCCT GAGTACGTGA ACCTCCCCAT CAATGGCAAC GGGAAACAGT
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721 AAATTTCTAA GAGAGCT

10
GENBANK ID: XM_016524.4
DEFINITION HOMO SAPIENS CREATINE KINASE, MITOCHONDRIAL 1 (UBIQUITOUS)
(CKMT1), MRNA.
VERSION XM_016524.4 GI:17477504
CDS 358..1704
/CODON_START=1

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1 CGCGCGAGTC TCAGGTCCCG CTAATTACCT GGCGGGTGCT GCGGCGGCTT GCGGCGGCTT
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121 GGCTGGGGCC TCCCTCTCCT CCGCCCCGCC GCCTGCCACT AGCTCATTCG GCCTCTCCTG
181 CAGTCTGATT GGCACCGGCT CCCATTCCGG CTCCAGCCTC CAATCCGACC CCCATTTCGG
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301 CCGGATCTTA TCTTGCGCCA GCGCCTACTC CAGGATCCCG TAGCCAGACC TCAAGCCATG
361 GCTGGTCCCT TCTCCCGTCT GCTGTCCGCC CGCCCGGGAC TCAGGCTCCT GGCTTTGGCC
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481 GAACGACGGA GGCTGTATCC CCCGAGCCAG ACATGGCCAA CTGGACAGCT CCCAGGTAAC
541 TGCACTAGGT CTAGGCGTCT GTGCCCTCCC TCCATGGTTA CTGGGTACCC CCTCCCCAGC
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601 GCTGAGTACC CAGACCTCCG AAAGCACAACT AACTGCATGG CCAGTCACCT GACCCAGCA
661 GTCTATGCAC GGCTCTGCGA CAAGACCACA CCCACTGGTT GGACGCTAGA TCAGTGTATC
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961 CGAGGACTCA GTCTGCCTCC AGCTTGCACT CGAGCAGAGC GACGAGAGGT GGAACGTGTT
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1201 TGGCACAACA ATGAGAAGAG CTTCCTGATC TGGGTGAATG AGGAGGATCA TACACGGGTG
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1501 CAAAAACGTG GTACTGGAGG AGTGGACACT GCTGCTACAG GCGGTGTCTT TGATATTTCT
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1681 CCTGTCATCC ACACCAAGCA TTAAGTCCCC ATCGCCAGCT GATGACTCAA GATTCCCAGG
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1801 TGCCTCCATC CTAGTAAAGA CTCCTTGCTA TGCTGC

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GENBANK ID: NM_001443.1
DEFINITION HOMO SAPIENS FATTY ACID BINDING PROTEIN 1, LIVER (FABP1), MRNA.
VERSION NM_001443.1 GI:4557576
CDS 43..426

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1 AGAGCCGCAG GTCAGTCGTG AAGAGGGAGC TCTATTGCCA CCATGAGTTT CTCCGGCAAG
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121 GAGCTCATCC AGAAGGGGAA GGATATCAAG GGGGTGTCGG AAATCGTGCA GAATGGGAAG
181 CACTTCAAGT TCACCATCAC CGCTGGGTCC AAAGTGATCC AAAACGAATT CACGGTGGGG
241 GAGGAATGTG AGCTGGAGAC AATGACAGGG GAGAAAGTCA AGACAGTGGT TCAGTTGGAA
301 GGTGACAATA AACTGGTGAC AACTTTCAA AACATCAAGT CTGTGACCGA ACTCAACGGC
361 GACATAATCA CCAATACCAT GACATTGGGT GACATTGTCT TCAAGAGAAT CAGCAAGAGA
421 ATTTAAACAA GTCTGCATTT CATATTATTT TAGTGTGTAA AATTAATGTA ATAAAGTGAA
60
481 CTTTGTGTTT

65
GENBANK ID: NM_001220.1
DEFINITION HOMO SAPIENS CALCIUM/CALMODULIN-DEPENDENT PROTEIN KINASE (CAM
KINASE) II BETA (CAMK2B), MRNA.
VERSION NM_001220.1 GI:10835005
CDS 47..1675

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1681 TGCGCCCTGG TTTCGCCGGA CAGAGTTGGT GTTTGGAGCC CGACTGCCCT CGGGCACACG
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1801 AAAACAAGAC CAGATGTGAT TTGTT

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GENBANK ID: NM_001677.1
 DEFINITION HOMO SAPIENS ATPASE, NA+/K+ TRANSPORTING, BETA 1 POLYPEPTIDE
 (ATP1B1), MRNA.
 VERSION NM_001677.1 GI:4502276
 CDS 127..1038

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1 GAATTCATGC TAAATTGCTG GAAGGCTGCG TCTCTGCTGT GGTGTCAATT CCGGATGCCT
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121 ATCGCCATGG CCCGCGGGAA AGCCAAGGAG GAGGGCAGCT GGAAGAAATT CATCTGGAAC
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1441 TATGTGAGCA AGGTTTGCTG TCCAAGGTGT AAATATTCAA CCGGAATAAA ACTGGCATGG
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5 1561 ATGAGCATT TTAACATACT CCATAGTCTT TTCCTGTGGT GTTAGGTCTT TATTTTTATT
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DEFINITION HOMO SAPIENS APOLIPOPROTEIN A-IV (APOA4), MRNA.

VERSION XM_052144.2 GI:15314431

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VERSION M29366.1 GI:181979
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40 GENBANK ID: L35572
VERSION L35572.1 GI:531219
MMVHCAGCERPIILDRFLNVLDRWHIKCVQCCECKTNLSEKCF
SREGKLYCKNDFRRFGTKCAGCAQGISPSDLVRKARSKVFHLNCFMVCNKQLSTG
45 EELYVIDENKFVCKDDYLSSSSSLKEGSLNSVSSCTDRSLSPDLQDPLQDDPKETDNST
SSDKETANNENEEQNSGTRRGPRRTTIKAKQLETAKAFAATPKPTRHIREQLAQETG
LNMRVIOVWFQNRSSKERRMKQLSALGARRHAFFRSPRRMRPLGGRLDESEMLGSTPY
TYYGDYQSDYAPGGNYDFFAHGPPSQASPADSSFLAASGPGSTPLGALEPPLAGPH
50 GADNPRFTDMISHPDTSPPEGLPGALHPMPGEVFSGGPSPPFPMSGTSGYSGPLSHP
NPENEAADV
GENBANK ID: X55787.1
VERSION X55787.1 GI:296022
MKAADVLDKPTLTIIKTEKVDLELFPSPEMECADVPLSTPSSKEM
MSQALKSYFSGFTKEQRRGIPKDPROWTDTHVRDWVMWAVNEFSLKGVDFHKFCMSG
AAVCALGKECFLELAPDFVGDILWEHLEILQKEDVKPYQVNGANPTYPESCYTSVYFI
55 SYGIEHAQCVPPEFSEPSFITESYQTLHPISSEELLSLKYENDYPSVILRDPLQET
LQTDYFRIKQEVLTNNMCLGRASRGKLGQDSFESVESYDSCDRLTQSWSSQSSFNS
60 LQRVPSYDSFDYEDYPAALPNHKPKGTFKDYVRDRADLNKDKPVIAPAAALAGYTGSGP
IQLWQFLLELLTDKSCQSFISWTGDCWEFKLSDPDEVARRWGRKNKPSMNYEKLRA
LRYYYDKNIIHKTAGNAYVYAFVCDLQSLGYPPEELHAMLDVKPDAD

65 GENBANK ID: M11507.1
DEFINITION Human transferrin receptor

VERSION M11507.1 GI:339515

MMDQARSAFSNLFGGEPLSYTRFSLARQVDGDNHVMKLAUDE
EENADNNTKANVTKPKRCSGSICYGTIAVIVFFLIGFMIGYLGKVEPKTECERLA
GTESPVREEPGEDFPAARRLYWDDLKRKLSEKLDSTDTSTIKLLNENSYVPREAGSQ
KDENLALYVENQFREFKLSKVWRDQHFVKIQVKDSAQNSVIVDKNGRLVYLVENPGG
YVAYSKAATVTGKLVHANFGTKKDFEDLYTPVNGSIVIVRAGKITFAEKVANAESLNA
IGVLIYMDQTKFPIVNAELSFEGHAHLGTGDPYTPGFPSFNHTQFPSPRSSGLPNIPV
QTISRAAAEKLFGNMEGDCPSDWKTDSTCRMVTSSEKNVKLTVSNVLKEIKILNIFGV
IKGFVEPDHYVVVGAQRDAWGPGAAKSGVGTALLKLAQMFSMDVLKDGFPQRSIIIF
ASWSAGDFGSGVATEWLEGYLSSLHLKAFTYINLDKAVLGTSNFKVSASPLLYTLIEK
TMQNVKHPVTGQFLYQDSNWASKVEKLTLDNAAFPFLAYSGIPAVSFCFCEDTDYPYL
GTTMDTYKELIERIPELNKVARAAAEVAGQFVIKLTVDVELNLDYERYNSQLLSFVRD
LNQYRADIKEMGLSLQWLYSARGDFFRATSRLTTDFGNAEKTDRFVMKKLNDVRMRVE
YHFLSPYVSPKESPFRRHVFWGSGSHTLPALLENLKLKQNNGAFNETLFRNQLALATW
TIQGAANALSGDVWDIDNEF

GENBANK ID: AAA60255

VERSION AAA60255.1 GI:190927

1 mteyklvvvg aggvvgksalt iqliqnhfvd eydptiedsy rkqvvidget clldildtag
61 qeysamrdq ymrtgegflc vfainnsksf adinlyreqi krvkdsddvp mvlvgnkcdl
121 ptrtvdtkqa helaksygip fietsaktrq gvedafytlv reirqymkk lnsddgtqg
181 cmglpcvwm

GENBANK ID: AAB38309

VERSION AAB38309.1 GI:1497931

1 mslvlnldlli ccrqlehdra terkkevekf krlirdpeti khldrhdsdk qgkylndwav
61 frflqkyiqk etecclriakp nvsastqasr qkkmqeissl vkyfikcanr raprlkcqel
121 lnyimdtvkd ssngaiygad csnilldkil svrkwyceis qqqwlelfsv yfrlylkpsq
181 dvhrvlvari ihavtkgccc qtdglnskfl dffskaiqca rkeksssgln hilaaltifl
241 ktavnfir vcelgdeilp tlliywtqhr lndslkevii elfqlqiyih hpkgaktqek
301 gayestkwrs ilynlydllv neishigsrg kyssgfrnia vkenlielma dichqvfned
361 trsleisqsy tttqressdy svpcrkkie lgwevikdhl qksqndfdlv pwlqiatqli
421 skypaslpnc elspllmils qlpqqrhge rtpyvlrcit evalcqdkrs nlessqksdl
481 lklwnkiwci tfrgisseqi qaenfgllga iiqgslvevd refwkltfsg acrpsscavc
541 cltlalttsi vpgtvkmgie qnmcevnrsf slkesimkw1 lfyqlegdle nstevppilh
601 snfphlvlek ilvsltmknc kaamnfqsv pecehhqkdk eelsfsevee lflqttfdkm
661 dfltivrecg iekhqssigf svhqnlkesl drcllglsq llnnyssait nsetlvrcsr
721 llvgvlgcyc ymgviaeeea ykselfqkak slmqcagesi tlfknktnee frigslnmm
781 qlctrclsnc tkkspnkias gfflrlltsk lmdiadick slasfikkpf drgevesmed
841 dtngnlmeve dqssmnlnd ypdssvsdan epgesqstg ainplaeeyl skqdllfldm
901 kflclcvtt aqntvsvfra adirrkllml idsstleptk slhlmym1 lkelppgeyp
961 lpmedvlell kplsnvcsly rrdqdvcti lnhvlhvkn lgqsnmdsen trdaqgqflt
1021 vigafwhltk erkyifsvrm alvnciktl1 eadpyskwai lnmvgkdfpv nevftqflad
1081 nhhqvrmlaa esinrlfqdt kgdssrllka lplklqqtat enaylkaqeg mremshsaen
1141 petldeiyar ksvlltliav vlscspicek qalfalcksv kenglephlv kkvekvset
1201 fgyrrledfm ashldylve wlnlqdeytn lssfpfilln ytniedfyr cykvliplhv
1261 irshfdevks ianqiqedwk slttdcfpki lvnilypyfay egtrdsqmaq qretatkvyd
1321 mlksenllgk qidhlfisnl peivvellmt lhpanssas qstdlcfsg dldpapnpph
1381 fpshvikatf ayisnchktk lksileilsk spdsyqkill aiceqaaetn nvykkhrilk
1441 iyhlfslll kdiksglgga wafvlrdviy tlihyinqrp scimdvslrs fslccdllsq
1501 vcqtavtyck dalenhlhvi vgtliplve qvevqkvld lkylvidnk dnenlyitik
1561 lldpfpdhv fkdrlritqk ikysrgpfs1 leehnhflsv svyda1plr leglkdlrrq
1621 lelhdqmv dmrassqnpq dgimvklvn llqlskmain htgekevlea vgsclgevgp
1681 idfstiaiqh skdasytka klfedkelqw tfimltylnn tlvedcvkvr saavtclkni
1741 latktghsfw eiymnttdpm laylqpfrts rkkflevprf dkenpfegld dinlwiplse
1801 nhdiwikltl cafldsggtk ceilqlkpm cevktdfcqt vlypylihdil lqdtnevrn
1861 llsthvqgff tsclrhfsqt srsttpanld sesehffrcc ldkksqrml avvdymrrqk
1921 rpssgtifnd afwldlnyle vakvaqcaa hftallyaei yadkksmddq ekrslafeeg
1981 sqsttissls ekskeetgis lqdlleiy1 sigepdslyg cgggkmlqpi trlrtyeha
2041 mwgkalvtyd letaipsstr qagiiqalqn lglchilsvy lkgldyenkd wcpelaelhy
2101 qaawrmqwd hctsvskeve gtsyheslyn alqslrdref stfyelskya rvkeveemck
2161 rslesvysly ptlsrlqag elesigelfs rsvthrlse vyikwqkhsq llkdsdfsfg
2221 epimalrtvi leilmekemd nsqrecikdi ltkhlvelsi lartfkntql peraifqikq

5 2281 ynsvscgvse wqleeaqvw akkeqslals ilkqmikkld ascaannpsl kltyteclrv
2341 cgnwlaetcl enpavimqty lekavevagn ydgessdelr ngkmkafsl arfsdtqyqr
2401 ienymkssef enkqallkra keevglireh kiqtnrytvk vqreleldel alralkedrk
2461 rflckaveny incllsgeeh dmwvfrlcsl wlensgvsev ngmmkrdgmk iptykflplm
2521 yqlaarmgtk mmggglgfhev lnnlisrism dhphtlfiil lalanandrde fltkpevarr
2581 sritknvpkq ssqldedrte aanriictir srpqrsvrsv ealcdayiil anldatqwt
2641 qrkqinipad qpitklknl dvvvpmeik vdhtgeygnl vtiqsfkaef rlaggvnlpk
2701 iidcvgsdsk errqlvkgrd dlrqdamvqq vfmcentllq rntetrkrkl tictykvvpl
2761 sqrsqglewc tgtvpigefl vnnedgahkr yrpndfsafq cqkkmmevqk ksfeekyevf
10 2821 mdvcqnfqpv fryfcmekfl dpaiwfekrl aytrsvatss ivgyilglgd rhvqnline
2881 qsaelvhidl gvafeqgkil ptpetvpfrr trdivdgmgi tgvegvrirc cektmevmrn
2941 sqetlltive vllypdplfdw tmnpkalyi qqrpedetel hptlnaddqe ckrnlsdidq
3001 sfdkvaervl mrlqekikgv eegtvlsvgg qvnlliqqai dpknlsrlfp gwkaw

15 //

20 GENBANK ID: AAA59145.1
VERSION AAA59145.1 GI:307058

1 mlkpslpfts llflqlpllg vglnttiltp ngnedttadf flttmptdsl svstlplpev
61 qcfvfnveym nctwnsssep qptnltlhyw yknsdndkvq kcshylfsee itsgcqlqkk
121 eihlyqtfvv qlqdprrrr qatqmkllqn lvipwapenl tlhklseql elnwnnrfln
181 hclehlvqyr tdwdhswteq svdyrhkfsi psvdgqkryt frvrsrfnpl cgsaqhwsew
25 241 shpihwgsnt skenpflfal eavvisvgsm gliisllcvy fwletmpri ptlknledlv
301 teyhgnfsaw sgvskglaes lqpdysrlc lvseippkkg algpggasp cnqhsqywap
361 pcytlkpet

30 GENBANK ID: AAC50825.1
VERSION AAC50825.1 GI:1117984

1 mvaprplrrv vlffyqgklcs magnfwqssh ylwildkqd llkerqkdlk flseeeywkl
61 qifftnviqa lgehlklrqq viatatvyfk rfyarysiks idpvlmaptc vflaskveef
35 121 gvsntria aatsvlktrf syafpkefpy rmnhilecef yllemddcl ivyhyprpl
181 qyvqdmged mllplawriv ndtyrtdlcl lyppfmiala clhvacvvqq kdarqwaef
241 svdmekilei irvilkllyeq wknfderkem atilskmpkp kpppnsegeq gpngsqnssy
301 sqs

40 GENBANK ID: AAC50473.1
VERSION AAC50473.1 GI:1314346

45 1 mamssggsgg gvpeqedsvl frrgtgqsd sdiwddtali kaydkavasf khalkngdic
61 etsgkpkttp krkpakknks qkntaaslq qwkvgdkcsa iwsedgciyp atiasidfr
121 etcvvytyg gnreeqnlsd llspicevan nieqnaqene nesqvstdes ensrspgnks
181 dnkpkpsapw nsflpppppm pgprlgpgkp glkfngpppp pppppphlls cwlppfsgp
50 241 piipppppic pdslddadl gsmliswys gyhtgyymgf rqnqkegrcs hsln

55 GENBANK ID: CAC15525
VERSION CAC15525.1 GI:11137517

1 mvsrdqahlg pkyvglwdfk srtdeelsfr agdvfhvark eeqwwatll deaggavagg
61 yvphnylaer etvesepwff gcisrseavr rlqaegnag aflirvsekp sadyvlsvrd
121 tqavrhykiw rraggrhlh eavsfslpe lvnyhraql shglrlaapc rkhepeplph
60 181 wddwerpree ftlcrklsg yfgevfeqlw kdrvqvaiqv isrdnllhqq mlqseiqamk
241 klrhkhilal yavvsvgdpv yiitelmakg sllellrdsd ekvlpvsell diawqvaegm
301 cylesqnyih rdlaarnilv gentlckvqd fglarliked vylshdnip ykwtapeals
361 rghystksdv wsfqillhem fsrgqvpyppg msnheafllv dagyrmpcpl ecppsvhklm
421 ltcwcrdpeq rpcfkalrer lssftsypn t

65

GENBANK ID: AAB84296
VERSION AAB84296.1 GI:2613135

5

1 tsttvrglna strylfrvra svqglgdwn tveettlglq saspvqesrv aedgldqqlv
61 lavvgsvsat cltilaalla lvcirrsclh rrhtftyqsg sgeetilqfs sgtltltrp
121 kpqpeplsyple

10

GENBANK ID: X63594.1
VERSION X63594.1 GI:57673
MFQFAGHGQDWAMEGPRDGLKKERLVDDRHDSGLDSMKDEDEYEQ
MVKELREIRLQPEAPLAAEPWKQQLTEDGDSFLHLAIHHEEKTILTMEVIGQVKGDLA
FLNFQNNLQQTPLHLAVITNQPGIAEALLKAGCDPELRDFRGNTPLHLACEQGCLASV
AVLTQTCTPQHLHSVLQATNYNGHTCLHLASIHGYLGIVEHLVTLGADVNAQEPENGR
TALHLAVDLQNPDLVSLLLKCGADVNRVTYQGYSPYQLTWGRPSTRIQQQLGQLTLEN
LQTLPESEDEESYDTESEFTEDELPYDDCVFGGQRLTL

15

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GENBANK ID: AAB60641

25

VERSION AAB60641.1 GI:516515

30

1 meqgdqsmke gmgttwllst pqhwlmqqfy netyygrtge fmedfpltl1 wsvtvsmpf
61 ggfigsllvg plvknfgrkg allfnnifsi vpailmgcsr vatsfeliii srllvgicag
121 vssnvvpmyl gelapknrg algvvpqlfi tvgilvaqif glrnllanvd gwpillgltg
181 vgaalql111 pffpespryl liqkkdeaaa kkalqtlrgw dsvdrevaai rgedeaekaa
241 gfigsvklfr mrsrlwqls iivlmggqql sgvnaiyya dqiylsagvp eehvgyvtg
301 tgavnvmtf cavfvvellg rrl11111gfs icliaccvlt aalalqdtvs wmpyisivcv
361 isyvighalg pspipallit eiflqssrps afmvggsvhw lsnftvglif pfigeglgpy
421 sfivfavicl lttiyifliv petkaktfie inqiftkmnk vsevypekee lkelpptse
481 q

35

GENBANK ID: M29069
VERSION M29069.1 GI:205553

5 MLSCCTTSTMPGMICKNSDLEFDSLKPCFYPEDDDIYFGGRNSTP
PGEDIWKKFELLPTPRLSPGRALAEDSLEPANWATEMLLPEADLWSNPAAEEDI FGLK
GLSGSSSNPVVLQDCMWSGFSREKPEVTVSEKLPGGCGSLAVGAGTLVPGAAAATSA
GHARSGTAGVGRKAAWLTELSHLDSECVDSAVIFPANKRESMPVATIPASAGAAISL
GDHQGLSSSLEDFLSNSGYVEEGEEIYVVMLGETQFSKTVTKLPTAAHSENAALTPE
10 CAQSGELILKRSDLIQEQHNYAAPPLPYAEDARPLKKPRSQDPLGPLKCVLRPKAPRL
RSRSNSDLEDIERRRNHNRMERQRRDIMRSSFLNLRDLVPELVHNEKAAKVVLKKAT
EYIHTLQTDSEKLLVEREKLYERKQQLLEKIKQSAVC

15 GENBANK ID: M29039.1
VERSION M29039.1 GI:186626

MCTKMEQPFYHDDSYTATGYGRAPGGLSLHDYKLLKPSLAVNLA
DPYRSLKAPGARGPGPEGGGGSGYFSGQSDTGASLKLASSELERLIVPNSNGVITTT
PTPPGQYFYPRGGGSGGAGGAGGVTEEQEGFADGFVKALDDLHKMNHVTPPNVSLG
20 ATGGPPAGPGGVYAGPEPPPVTNLSSYSPASASSGGAGAAGVTGSSYPTTTISYLPH
APPFAGGHPAQLGLGRGASTFKEEPQTVPEARSRDATPPVSPINMEDQERIKVERKRL
RNRLAATKCRKRKLERIARLEDKVTKLKAENAGLSSTAGLLREQVAQLKQKVMTHVSN
GCQLLLGVKGHAF

25 GENBANK ID: X56681.1
VERSION X56681.1 GI:34018

30 METPFYGDDEALSGLGGGASGSGGTFA SPGRLEFGAPPTAAAGSM
MKKDALTLSEQLVAAALKPAPAPASYPPAADGAPSAAPPDGLLASPDGLLKLASPE
LERLIQSNGLVTTTPTSSQFLYPKVAASEEQEFAEGFVKALEDLHKQNLGAGRAAA
AAAAAAGGPGSGTATGSAPPGLAPAAAPEAPVYANLSSYAGGAGGAGGAATVAFAAE
PVPFPPPPPPGALGPRLAALKDEPQTVDPVPSFGESPPLSPIDMDTQERIKAEKRL
35 RNRIAASKCRKRKLERISRLEEKVKTLKSQNTLASTASLLREQVAQLKQKVLSHVNS
GCQLLPQHQPAY

40 GENBANK ID: NM_003150.1
VERSION NM_003150.1 GI:4507252

MAQWNQLQQLDTRYLEQLHQLYSDSFPMELRQFLAPWIESQDWA
YAASKESHATLVFHNLLGEIDQYSRFLQESNVLYQHNLRRIKQFLQSRYLEKPMETIA
RIVARCLWEESRLLQTAATAAQGGQANHPTAAVVTEKQOMLEQLQDVRKRVQDLEQ
45 KMKVVENLQDDDFDNKYTLKSQGMQDLNNGNSVTRQKMQLEQMLTALDQMRRSIV
SELAGLLSAMEYVQKTLTDEELADWKRQQIACIGGPPNICLDRLENWITSLAESQLQ
TRQIKKLEELHQKVSYGDPVQHRPMLERIVELFRNLMKSAFVVERQPCMPMHPD
RPLVIKTGVQFTTKVRLLVKFPNELNYQLKIKVCIDKDSGDVAALRGSRKFNILGTNTK
VMNMEESNNGSLSAEFKHLTLREQRCGNGGRANCDA SLIVTEELHLITFETEVYHQGL
50 KIDLETHSLSVVISNICQMPNAWASILWYNMLTNNPKNVNFFTKPPIGTWDQVAEVL
SWQFSSTTKRGLSIEQLTTAEKLLGPGVNYSGCQITWANFCKENMAGKGFSYVWVLD
NIIDLVKKYILALWNEGYIMGFISKERERAILSTKPPGTFLRFSESSKEGGVTFTWV
EKDISGKTQIQSVEPYTKQQLNMSFAEIIIMGYKIMDATNILLSPLVLYPDIPKEEA
FGKYCRPESQEHPEADPGSAAPYLKTKFICVTPTTCSNTIDLPMSPRALDSLMQFGNN
55 GEGAEPGAGGQFESLTFDMELTSECATSPM

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GETING PEYER'S PATCHES AND M CELL RECEPTORS

(57) Abstract: Methods of increasing of or decreasing the levels of a protein in a PP cell; methods of increasing antigen, vaccine,
DNA vaccine delivery to M cells, use of human serum albumin and other transport enhancing proteins to enhance oral drug delivery;
use of calreticulin to enhance oral antigen delivery, use of other cell surface proteins, receptors, and transporters to enhance delivery
to M cells of antigens or vaccine delivery vehicles, use of other cytoplasmic proteins to regulate intracellular trafficking and delivery
to mucosal immune sampling and processing systems.

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International Application No
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A61K39/00 A61K47/48

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

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Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the International search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, MEDLINE, EMBASE, BIOSIS

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>US 6 060 082 A (CHEN HONGMING ET AL) 9 May 2000 (2000-05-09) column 1, line 11 - line 26 column 2, line 38 - line 47 column 2, line 66 - column 3, line 44 column 7, line 4 - line 33 column 19, line 31 - column 22, line 35 ----- -/--</p>	1-58, 61-64

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☒ Patent family members are listed in annex.

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- "&" document member of the same patent family

Date of the actual completion of the international search

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Date of mailing of the international search report

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International Application No

PCT/IB 02/03866

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X	GULLBERG ELISABET ET AL: "Expression of specific markers and particle transport in a new human intestinal M-cell model." BIOCHEMICAL AND BIOPHYSICAL RESEARCH COMMUNICATIONS, vol. 279, no. 3, 29 December 2000 (2000-12-29), pages 808-813, XP002252802 ISSN: 0006-291X page 808 abstract page 810, right-hand column, paragraph 1 -----	59,60
A	US 6 117 632 A (O'MAHONY DANIEL JOSEPH) 12 September 2000 (2000-09-12) column 2, line 12 - line 40 column 3, line 48 - line 64 -----	59,60
A	HADDAD A ET AL: "TARGETED M CELL IMMUNIZATION FOR HIV-1 ENV DNA VACCINES" FASEB JOURNAL, FED. OF AMERICAN SOC. FOR EXPERIMENTAL BIOLOGY, BETHESDA, MD, US, vol. 14, no. 6, 20 April 2000 (2000-04-20), page A1204, XP000995418 ISSN: 0892-6638 abstract 185.7 -----	
E	WO 02/080852 A (O'MAHONY DANIEL J ;BRAYDEN DAVID J (IE); BYRNE DARAGH (IE); DIGITA) 17 October 2002 (2002-10-17) the whole document -----	1-58, 61-64

INTERNATIONAL SEARCH REPORT

International application No.
PCT/IB 02/03866

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.:
because they relate to subject matter not required to be searched by this Authority, namely:

Although partially claims 1-9, 12-21, 34-51, 61-64, and completely claims 10, 28-31, 33, are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. ☒ Claims Nos.: 1-58, 61-64 partially
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
see FURTHER INFORMATION sheet PCT/ISA/210
3. ☐ Claims Nos.:
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

see additional sheet

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☒ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

1-64

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
- ☐ No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box 1.2

Claims Nos.: 1-58, 61-64 partially

Present claims 1-58 and 61-64 relate to an extremely large number of possible compounds and the use thereof. In the present particular case, there is considered to be support within the meaning of Article 6 PCT and/or disclosure within the meaning of Article 5 PCT for none of the compounds comprised in the claims. More particularly, there is a complete lack of evidence that the compounds recited in the claims provide any plausible solution to a technical problem possibly addressed by the application. Consequently, the claims are considered to so lack support, and the application to so lack disclosure, that a meaningful search over the whole of the claimed scope is impossible. As a result, the search has been carried out for those elements of the application which appear to be supported and disclosed, and for which a meaningful search was considered possible: in this particular case, a search was only considered possible on the inventive concept underlying the application, ie essentially that related to the correlation of altered expression of proteins in Peyer's patch cells / M cells with their use in enabling / facilitating oral delivery of drugs and antigens.

The applicant's attention is drawn to the fact that claims relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure. If the application proceeds into the regional phase before the EPO, the applicant is reminded that a search may be carried out during examination before the EPO (see EPO Guideline C-VI, 8.5), should the problems which led to the Article 17(2) declaration be overcome.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-64

Means to increase in PP cells of the intestine levels of proteins specific to PP cells, or decrease in PP cells of the intestine the levels of proteins non-specific to PP cells, via delivery of a nucleic acid encoding the PP cell specific protein / the protein per se, or delivery of an antisense / ribozyme / RNAi to the protein non-specific to PP cells, (eg claim 1 et seq., claim 15 and seq.); means to deliver a composition to a PP cell wherein said composition has a ligand that will specifically bind to a PP specific protein (eg claim 57); related subject matter

2. claim: 65

Promote enterocyte-M cell conversion via use of an antigen and a bacteria, probiotic yoghurt or bacterial component

INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

PCT/IB 02/03866

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